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Welcome to STN International NEWS Web Page URLs for STN Seminar Schedule - N. America NEWS "Ask CAS" for self-help around the clock NEWS 3 DEC 23 New IPC8 SEARCH, DISPLAY, and SELECT fields in USPATFULL/ USPAT2 NEWS JAN 13 IPC 8 searching in IFIPAT, IFIUDB, and IFICDB NEWS 5 JAN 13 New IPC 8 SEARCH, DISPLAY, and SELECT enhancements added to NEWS 6 JAN 17 Pre-1988 INPI data added to MARPAT IPC 8 in the WPI family of databases including WPIFV NEWS 7 JAN 17 NEWS 8 JAN 30 Saved answer limit increased NEWS 9 FEB 21 STN AnaVist, Version 1.1, lets you share your STN AnaVist visualization results NEWS 10 FEB 22 The IPC thesaurus added to additional patent databases on STN NEWS 11 FEB 22 Updates in EPFULL; IPC 8 enhancements added NEWS 12 FEB 27 New STN AnaVist pricing effective March 1, 2006 NEWS 13 FEB 28 MEDLINE/LMEDLINE reload improves functionality NEWS 14 FEB 28 TOXCENTER reloaded with enhancements NEWS 15 FEB 28 REGISTRY/ZREGISTRY enhanced with more experimental spectral property data NEWS 16 MAR 01 INSPEC reloaded and enhanced NEWS 17 MAR 03 Updates in PATDPA; addition of IPC 8 data without attributes NEWS 18 MAR 08 X.25 communication option no longer available after June 2006 NEWS 19 MAR 22 EMBASE is now updated on a daily basis NEWS 20 APR 03 New IPC 8 fields and IPC thesaurus added to PATDPAFULL NEWS 21 APR 03 Bibliographic data updates resume; new IPC 8 fields and IPC thesaurus added in PCTFULL NEWS 22 APR 04 STN AnaVist \$500 visualization usage credit offered NEWS 23 APR 12 LINSPEC, learning database for INSPEC, reloaded and enhanced NEWS 24 APR 12 Improved structure highlighting in FQHIT and QHIT display in MARPAT NEWS 25 APR 12 Derwent World Patents Index to be reloaded and enhanced during second quarter; strategies may be affected

NEWS EXPRESS FEBRUARY 15 CURRENT VERSION FOR WINDOWS IS V8.01a,
CURRENT MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP),
AND CURRENT DISCOVER FILE IS DATED 19 DECEMBER 2005.
V8.0 AND V8.01 USERS CAN OBTAIN THE UPGRADE TO V8.01a AT
http://download.cas.org/express/v8.0-Discover/

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=> ile registry

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=> file registry

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SINCE FILE TOTAL ENTRY SESSION 0.21 0.21

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STRUCTURE FILE UPDATES: 24 APR 2006 HIGHEST RN 881733-90-0 DICTIONARY FILE UPDATES: 24 APR 2006 HIGHEST RN 881733-90-0

New CAS Information Use Policies, enter HELP USAGETERMS for details.

TSCA INFORMATION NOW CURRENT THROUGH January 6, 2006

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REGISTRY includes numerically searchable data for experimental and predicted properties as well as tags indicating availability of experimental property data in the original document. For information on property searching in REGISTRY, refer to:

http://www.cas.org/ONLINE/UG/regprops.html

Uploading C:\Program Files\Stnexp\Queries\10619662query.str

chain nodes :

7 8 9 10 11 12 13 14 15 16 17 18 19 20 21 22 23

ring nodes :

1 2 3 4 5 6 chain bonds:

1-23 2-7 6-8 7-10 7-11 8-9 8-12 11-13 11-16 12-14 12-15 15-17 15-21

15-22 16-18 16-19 16-20

ring bonds :

1-2 1-6 2-3 3-4 4-5 5-6

exact/norm bonds :

1-2 1-6 2-3 3-4 4-5 5-6 7-10 7-11 8-9 8-12 11-16 12-15 15-17 16-18

exact bonds :

1-23 2-7 6-8 11-13 12-14 15-21 15-22 16-19 16-20

Match level :

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:CLASS 8:CLASS 9:CLASS 10:CLASS 11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 16:CLASS 17:Atom 18:Atom 19:CLASS 20:CLASS 21:CLASS 22:CLASS 23:CLASS

9006

2 ANSWERS

L1 STRUCTURE UPLOADED

=> s L1

SAMPLE SEARCH INITIATED 11:50:07 FILE 'REGISTRY'
SAMPLE SCREEN SEARCH COMPLETED - 391 TO ITERATE

100.0% PROCESSED 391 ITERATIONS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**

BATCH **COMPLETE**

PROJECTED ITERATIONS: 6634 TO

PROJECTED ANSWERS: 2 TO 124

L2 2 SEA SSS SAM L1

=> d L2 1-2

L2 ANSWER 1 OF 2 REGISTRY COPYRIGHT 2006 ACS on STN

RN 544678-73-1 REGISTRY

ED Entered STN: 09 Jul 2003

CN 2,4-Pyridinedicarboxamide, N,N'-bis[(3-methylphenyl)methyl]-,
 monohydrochloride (9CI) (CA INDEX NAME)

MF C23 H23 N3 O2 . C1 H

SR CA

LC STN Files: CA, CAPLUS, USPAT2, USPATFULL

CRN (747406-86-6)

● HCl

- 1 REFERENCES IN FILE CA (1907 TO DATE)
- 1 REFERENCES IN FILE CAPLUS (1907 TO DATE)
- L2 ANSWER 2 OF 2 REGISTRY COPYRIGHT 2006 ACS on STN
- RN 449734-45-6 REGISTRY
- ED Entered STN: 12 Sep 2002
- CN 2,4-Pyridinedicarboxamide, N,N'-bis([1,1'-biphenyl]-2-ylmethyl)- (9CI) (CA INDEX NAME)

OTHER NAMES:

- CN Pyridine-2,4-dicarboxylic acid bis[[(biphenyl)-2-ylmethyl]amide]
- FS 3D CONCORD
- MF C33 H27 N3 O2
- SR CA
- LC STN Files: CA, CAPLUS, TOXCENTER, USPAT2, USPATFULL

$$\begin{array}{c|c} CH_2-NH-C & N & O \\ \hline \\ Ph & C-NH-CH_2 \\ \hline \\ Ph & Ph \\ \end{array}$$

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

- 1 REFERENCES IN FILE CA (1907 TO DATE)
- 1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

=> sel L2

E1 THROUGH E3 ASSIGNED

=> file caplus medline biosis
COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 5.34 5.55

FULL ESTIMATED COST

FILE 'CAPLUS' ENTERED AT 11:50:42 ON 25 APR 2006 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2006 AMERICAN CHEMICAL SOCIETY (ACS)

FILE 'MEDLINE' ENTERED AT 11:50:42 ON 25 APR 2006

FILE 'BIOSIS' ENTERED AT 11:50:42 ON 25 APR 2006 Copyright (c) 2006 The Thomson Corporation

=> s E1-E3

L3

2 ("PYRIDINE-2,4-DICARBOXYLIC ACID BIS(((BIPHENYL)-2-YLMETHYL)AMID E)"/BI OR 449734-45-6/BI OR 544678-73-1/BI)

=> d L3 1-2 ti abs bib

L3 ANSWER 1 OF 2 CAPLUS COPYRIGHT 2006 ACS on STN

TI Preparation of 2,4-pyridinedicarboxamides and 4,6-pyrimidinedicarboxamides as inhibitors of collagenase (MMP 13)
GI

$$R^1$$
 R^2
 R^3
 R^3
 R^2
 R^3

AB Title compds. [I; A = CH, N; R1-R3 = H, halo, (halogenated) alkyl, alkoxy, OH, CO2R4, cyano, NR5R6, etc.; R4 = H, alkyl; R5, R6 = H, alkyl, alkylcarbonyl, etc.; or R1R2, R2R3 = 5-6 membered (aromatic) (saturated) (hetero)cyclyl], were prepd for the treatment of degenerative joint diseases. Thus, 4,6-pyrimidinedicarboxylic acid in SOCl2 was stirred for 2 h at 85° followed by addition of CH2Cl2 at room temperature and Et3N at 0°. The reaction mixture was further stirred with 3-chloro-4-fluorobenzylamine for 15 min to give 40% N,N-bis(3-chloro-4-fluorobenzyl)pyrimidine-4,6-dicarboxamide. The latter inhibited collagenase 3 (MMP 13) with IC50 = 23 nM.

Ι

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AN 2003:467290 CAPLUS

DN 139:53028

TI Preparation of 2,4-pyridinedicarboxamides and 4,6-pyrimidinedicarboxamides as inhibitors of collagenase (MMP 13)

IN Habermann, Joerg; Weithmann, Klaus-Ulrich; Kogler, Herbert; Kirsch, Reinhard; Wehner, Volkmar

PA Aventis Pharma Deutschland G.m.b.H., Germany

SO Ger. Offen., 20 pp.

CODEN: GWXXBX

DT Patent

LA German

FAN.CNT 1

ran.			TENT NO. 				D	DATE			APPL	ICAT:	ION I	NO.		D	ATE	
ΡI	DE	1016	0357			A1		2003	0618	1	DE 2	001-	1016	0357		2	00112	208
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	US	6933	298			B2		2005	0823									
PRAI	DE	2001	-101	6035	7	Α		2001	1208									
	US	2002	-358	887P		P		2002	0222									
	WO	3 2002-358887P 3 2002-EP13240				W		2002	1125									

L3 ANSWER 2 OF 2 CAPLUS COPYRIGHT 2006 ACS on STN

TI Preparation of pyridinedicarboxamide and -dicarboxylic acid derivatives as selective MMP-13 matrix metalloproteinase inhibitors with therapeutic uses

GI

AΒ Selective MMP-13 inhibitors are pyridine derivs. (I; e.g. pyridine-2,4-dicarboxylic acid bis(3-methoxybenzylamide)) or a pharmaceutically acceptable salt thereof, wherein: R1 and R2 independently are H, halo, hydroxy, C1-C6 alkyl, C1-C6 alkoxy, C2-C6 alkenyl, C2-C6 alkynyl, NO2, NR4R5, CN, or CF3; E is independently O or S; A and B independently are OR4 or NR4R5; R4 and R5 independently are H, C1-C6 alkyl, C2-C6 alkenyl, C2-C6 alkynyl, (CH2)n aryl, (CH2)n cycloalkyl, (CH2)n heteroaryl, or R4 and R5 when taken together with the N to which they are attached complete a 3- to 8-membered ring containing C atoms and optionally containing a heteroatom selected from O, S, or NH, and optionally substituted or unsubstituted; n is 0 to 6. Although I and other Markush structures in the patent show 2,4- derivs., many specific 3,5- derivs. are included in the claims and examples. Combinatorial and non-combinatorial methods were used to prepare numerous claimed compds. and characterization data is reported for about 90 compds. IC50 values for various claimed compds. show the selectivity towards MMP-13 vs. MMP-1 and MMP-3 and the potent MMP-13 inhibitory activity (e.g. 0.033 µM for pyridine-2,4-dicarboxylic acid bis[((1,3-benzodioxol-5-yl)methyl)amide]).

AN 2002:637657 CAPLUS

DN 137:185420

TI Preparation of pyridinedicarboxamide and -dicarboxylic acid derivatives as selective MMP-13 matrix metalloproteinase inhibitors with therapeutic uses

IN Barvian, Nicole Chantel; Connor, David Thomas; O'brien, Patrick Michael;
Ortwine, Daniel Fred; Patt, William Chester; Shuler, Kevon Ray; Wilson,
Michael William

PA Warner-Lambert Company, USA

SO PCT Int. Appl., 68 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

	PATENT NO.					KIND DATE		APPLICATION NO.					DATE					
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			LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NO,	NZ,	OM,	PH,
			PL,	PT,	RO,	RU,	SD,	SE,	SG,	SI,	SK,	SL,	TJ,	TM,	TR,	TT,	TZ,	UA,
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			CY,	DE,	DK,	ES,	FI,	FR,	GB,	GR,	ΙE,	IT,	LU,	MC,	NL,	PT,	SE,	TR,
			BF,	ВJ,	CF,	CG,	CI,	CM,	GA,	GN,	GQ,	GW,	ML,	MR,	ΝE,	SN,	TD,	TG
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20020204
      EE 200300391 A 20031215 EE 2003-391
                                                   BR 2002-7863
BR 2002007863 A 20040427 BR 2002-7863
JP 2004529878 T2 20040930 JP 2002-564501
CN 1537101 A 20041013 CN 2002-804945
US 2002161000 A1 20021031 US 2002-71073
US 6881743 B2 20050419
ZA 2003006041 A 20041105 ZA 2003-6041
NO 2003003570 A 20030812 NO 2003-3570
BG 108089 A 20050131 BG 2003-108089
US 2004209922 A1 20041021 US 2004-842863
US 7015237 B2 20060321

PRAI US 2001-268781P P 20010214
WO 2002-IB345 W 20020204
US 2002-71073 A3 20020208
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      BR 2002007863
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20030813
20040510
      MARPAT 137:185420
                 THERE ARE 10 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT 10
                 ALL CITATIONS AVAILABLE IN THE RE FORMAT
=> file uspatfull
                                                           SINCE FILE
COST IN U.S. DOLLARS
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FULL ESTIMATED COST
                                                                  31.06
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CA SUBSCRIBER PRICE
FILE 'USPATFULL' ENTERED AT 11:52:38 ON 25 APR 2006
CA INDEXING COPYRIGHT (C) 2006 AMERICAN CHEMICAL SOCIETY (ACS)
FILE COVERS 1971 TO PATENT PUBLICATION DATE: 20 Apr 2006 (20060420/PD)
FILE LAST UPDATED: 20 Apr 2006 (20060420/ED)
HIGHEST GRANTED PATENT NUMBER: US7032245
HIGHEST APPLICATION PUBLICATION NUMBER: US2006085880
CA INDEXING IS CURRENT THROUGH 20 Apr 2006 (20060420/UPCA)
ISSUE CLASS FIELDS (/INCL) CURRENT THROUGH: 20 Apr 2006 (20060420/PD)
REVISED CLASS FIELDS (/NCL) LAST RELOADED: Feb 2006
USPTO MANUAL OF CLASSIFICATIONS THESAURUS ISSUE DATE: Feb 2006
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          144459 "PYRIDINE"/BI
         4429129 "2"/BI
         4369219 "4"/BI
           72135 "DICARBOXYLIC"/BI
          840420 "ACID"/BI
          221352 "BIS"/BI
           41674 "BIPHENYL"/BI
         4429129 "2"/BI
           12843 "YLMETHYL"/BI
          153431 "AMIDE"/BI
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                     BIPHENYL" (W) "2" (W) "YLMETHYL" (W) "AMIDE") /BI)
                0 449734-45-6/BI
                0 544678-73-1/BI
                4 ("PYRIDINE-2,4-DICARBOXYLIC ACID BIS(((BIPHENYL)-2-YLMETHYL)AMID
L4
                  E) "/BI OR 449734-45-6/BI OR 544678-73-1/BI)
=> d L4 1-4 ti abs bib
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ANSWER 1 OF 4 USPATFULL on STN

1.4

IE, SI, LT, LV, FI, RO, MK, CY, AL, TR

Pyridine matrix metalloproteinase inhibitors ΤI Selective MMP-13 inhibitors are pyridine derivatives of the formula ##STR1## or a pharmaceutically acceptable salt thereof, wherein: R.sup.1 and R.sup.2 independently are hydrogen, halo, hydroxy, C.sub.1-C.sub.6 alkyl, C.sub.1-C.sub.6 alkoxy, C.sub.2-C.sub.6 alkenyl, C.sub.2-C.sub.6 alkynyl, N0.sub.2, NR.sup.4R.sup.5, CN, or CF.sub.3; E is independently 0 or S; A and B independently are OR.sup.4 or NR.sup.4R.sup.5; R.sup.4 and R.sup.5 independently are H, C.sub.1-C.sub.6 alkyl, C.sub.2-C.sub.6 alkenyl, C.sub.2-C.sub.6 alkynyl, (CH.sub.2).sub.n aryl, (CH.sub.2).sub.n cycloalkyl, (CH.sub.2).sub.n heteroaryl, or R.sup.4 and R.sup.5 when taken together with the nitrogen to which they are attached complete a 3- to 8-membered ring containing carbon atoms and optionally containing a heteroatom selected from O, S, or NH, and optionally substituted or unsubstituted; n is an integer of from 0 to 6. CAS INDEXING IS AVAILABLE FOR THIS PATENT. 2004:268379 USPATFULL ΤI Pyridine matrix metalloproteinase inhibitors ΙN Barvian, Nicole Chantel, Ann Arbor, MI, UNITED STATES Connor, David Thomas, Ann Arbor, MI, UNITED STATES O'Brien, Patrick Michael, Stockbridge, MI, UNITED STATES Ortwine, Daniel Fred, Saline, MI, UNITED STATES Patt, William Chester, Chelsea, MI, UNITED STATES Wilson, Michael William, Ann Arbor, MI, UNITED STATES PΙ US 2004209922 **A1** 20041021 US 7015237 B2 20060321 ΑI US 2004-842863 **A1** 20040510 (10) Division of Ser. No. US 2002-71073, filed on 8 Feb 2002, PENDING RLIPRAI US 2001-268781P 20010214 (60) DT Utility FS APPLICATION LREP WARNER-LAMBERT COMPANY, 2800 PLYMOUTH RD, ANN ARBOR, MI, 48105 CLMN Number of Claims: 13 ECL Exemplary Claim: CLM-001-9 DRWN No Drawings LN.CNT 1657 CAS INDEXING IS AVAILABLE FOR THIS PATENT. L4 ANSWER 2 OF 4 USPATFULL on STN ΤI Combination of an allosteric carboxylic inhibitor of matrix metalloproteinase-13 with a selective inhibitor of cyclooxygenase-2 that is not celecoxib or valdecoxib AB This invention provides a combination, comprising an allosteric carboxylic inhibitor of MMP-13, or a pharmaceutically acceptable salt thereof, with a selective inhibitor of COX-2, or a pharmaceutically

This invention provides a combination, comprising an allosteric carboxylic inhibitor of MMP-13, or a pharmaceutically acceptable salt thereof, with a selective inhibitor of COX-2, or a pharmaceutically acceptable salt thereof, that is not celecoxib or valdecoxib. This invention also provides a method of treating a disease that is responsive to inhibition of MMP-13 and cyclooxygenase-2, comprising administering to a patient suffering from such a disease the invention combination comprising an allosteric carboxylic inhibitor of MMP-13, or a pharmaceutically acceptable salt thereof, with a selective inhibitor of COX-2, or a pharmaceutically acceptable salt thereof, that is not celecoxib or valdecoxib. This invention also provides a pharmaceutical composition, comprising the invention combination comprising an

allosteric carboxylic inhibitor of MMP-13, or a pharmaceutically acceptable salt thereof, with a selective inhibitor of COX-2, or a pharmaceutically acceptable salt thereof, that is not celecoxib or valdecoxib, and a pharmaceutically acceptable carrier, diluent, or excipient. This invention also provides a combination comprising an NSAID, or a pharmaceutically acceptable salt thereof, and an allosteric carboxylic inhibitor of MMP-13, or a pharmaceutically acceptable salt thereof. This invention also provides a pharmaceutical composition, comprising the invention combination comprising an allosteric carboxylic inhibitor of MMP-13, or a pharmaceutically acceptable salt thereof, with an NSAID, or a pharmaceutically acceptable salt thereof, and a pharmaceutically acceptable carrier, diluent, or excipient. This invention also provides a method of treating a disease that is responsive to inhibition of MMP-13 and cyclooxygenase-1 or cyclooxygenase-2, comprising administering to a patient suffering from such a disease the invention combination comprising an allosteric carboxylic inhibitor of MMP-13, or a pharmaceutically acceptable salt thereof, with an NSAID, or a pharmaceutically acceptable salt thereof. The invention combinations may also be further combined with other pharmaceutical agents depending on the disease being treated.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AN 2004:25212 USPATFULL

TI Combination of an allosteric carboxylic inhibitor of matrix metalloproteinase-13 with a selective inhibitor of cyclooxygenase-2 that is not celecoxib or valdecoxib

IN Roark, William Howard, Ann Arbor, MI, UNITED STATES

PI US 2004019054 A1 20040129

AI US 2003-619769 A1 20030715 (10)

PRAI US 2002-396785P 20020717 (60)

DT Utility

FS APPLICATION

LREP WARNER-LAMBERT COMPANY, 2800 PLYMOUTH RD, ANN ARBOR, MI, 48105

CLMN Number of Claims: 9

ECL Exemplary Claim: 1

DRWN No Drawings

LN.CNT 8368

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

- L4 ANSWER 3 OF 4 USPATFULL on STN
- TI Combination of an allosteric carboxylic inhibitor of matrix metalloproteinase-13 with celecoxib or valdecoxib
- This invention provides a combination, comprising an allosteric AB carboxylic inhibitor of MMP-13, or a pharmaceutically acceptable salt thereof, with celecoxib, or a pharmaceutically acceptable salt thereof, or valdecoxib, or a pharmaceutically acceptable salt thereof. This invention also provides a method of treating a disease that is responsive to inhibition of MMP-13 and cyclooxygenase-2, comprising administering to a patient suffering from such a disease the invention combination comprising an allosteric carboxylic inhibitor of MMP-13, or a pharmaceutically acceptable salt thereof, with celecoxib, or a pharmaceutically acceptable salt thereof, or valdecoxib, or a pharmaceutically acceptable salt thereof. This invention also provides a pharmaceutical composition, comprising the invention combination comprising an allosteric carboxylic inhibitor of MMP-13, or a pharmaceutically acceptable salt thereof, with celecoxib, or a pharmaceutically acceptable salt thereof, or valdecoxib, or a pharmaceutically acceptable salt thereof, and a pharmaceutically acceptable carrier, diluent, or excipient.

The invention combination may also be further combined with other pharmaceutical agents depending on the disease being treated.

CAS INDEXING IS AVAILABLE FOR THIS PATENT. AN 2004:25211 USPATFULL

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Combination of an allosteric carboxylic inhibitor of matrix
      metalloproteinase-13 with celecoxib or valdecoxib
IN
      Roark, William Howard, Ann Arbor, MI, UNITED STATES
      US 2004019053
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                         A1
                               20040129
      US 2003-619662
AΙ
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                               20030715 (10)
PRAI
      US 2002-396903P
                          20020717 (60)
DT
      Utility
FS
      APPLICATION
      WARNER-LAMBERT COMPANY, 2800 PLYMOUTH RD, ANN ARBOR, MI, 48105
LREP
CLMN
      Number of Claims: 9
ECL
      Exemplary Claim: 1
DRWN
      No Drawings
LN.CNT 8040
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
    ANSWER 4 OF 4 USPATFULL on STN
L4
       Pyridine matrix metalloproteinase inhibitors
TI
AB
      Selective MMP-13 inhibitors are pyridine derivatives of the formula
      ##STR1##
      or a pharmaceutically acceptable salt thereof,
      wherein:
      R.sup.1 and R.sup.2 independently are hydrogen, halo, hydroxy,
      C.sub.1-C.sub.6 alkyl, C.sub.1-C.sub.6 alkoxy, C.sub.2-C.sub.6 alkenyl,
      C.sub.2-C.sub.6 alkynyl, NO.sup.2, NR.sup.4R.sup.5, CN, or CF.sub.3,
      E is independently O or S;
      A and B independently are OR.sup.4 or NR.sup.4R.sup.5;
       R.sup.4 and R.sup.5 independently are H. C.sub.1-C.sub.6 alkyl,
       C.sub.2-C.sub.6 alkenyl, C.sub.2-C.sub.6 alkynyl, (CH.sub.2).sub.n aryl,
       (CH.sub.2).sub.n cycloalkyl, (CH.sub.2).sub.n heteroaryl, or R.sup.4 and
       R.sup.5 when taken together with the nitrogen to which they are attached
       complete a 3 to 8-membered ring containing carbon atoms and optionally
       containing a heteroatom selected from O, S, or NH, and optionally
       substituted or unsubstituted,
       n is an integer of from 0 to 6.
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       2002:288130 USPATFULL
       Pyridine matrix metalloproteinase inhibitors
TI
IN
       Barvian, Nicole Chantel, Ann Arbor, MI, UNITED STATES
       Connor, David Thomas, Ann Arbor, MI, UNITED STATES
       O'Brien, Patrick Michael, Stockbridge, MI, UNITED STATES
       Ortwine, Daniel Fred, Saline, MI, UNITED STATES
       Patt, William Chester, Chelsea, MI, UNITED STATES
       Shuler, Kevon Ray, Chelsea, MI, UNITED STATES
       Wilson, Michael William, Ann Arbor, MI, UNITED STATES
PΙ
      US 2002161000
                          A1
                               20021031
      US 6881743
                          B2
                               20050419
ΑI
      US 2002-71073
                          A1
                               20020208 (10)
PRAI
      US 2001-268781P
                           20010214 (60)
DT
       Utility
       APPLICATION
FS
       Claude F. Purchase, Jr., Warner-Lambert Company, 2800 Plymouth Road, Ann
LREP
       Arbor, MI, 48105
CLMN
      Number of Claims: 35
ECL
       Exemplary Claim: 1
       No Drawings
DRWN
LN.CNT 1991
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
```

ΤI

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=> file uspgpubs
'USPGPUBS' IS NOT A VALID FILE NAME
SESSION CONTINUES IN FILE 'USPATFULL'
Enter "HELP FILE NAMES" at an arrow prompt (=>) for a list of files
that are available. If you have requested multiple files, you can
specify a corrected file name or you can enter "IGNORE" to continue
accessing the remaining file names entered.
=> file pgpub
'PGPUB' IS NOT A VALID FILE NAME
SESSION CONTINUES IN FILE 'USPATFULL'
Enter "HELP FILE NAMES" at an arrow prompt (=>) for a list of files
that are available. If you have requested multiple files, you can
specify a corrected file name or you can enter "IGNORE" to continue
accessing the remaining file names entered.
=> file uspgpub
'USPGPUB' IS NOT A VALID FILE NAME
SESSION CONTINUES IN FILE 'USPATFULL'
Enter "HELP FILE NAMES" at an arrow prompt (=>) for a list of files
that are available. If you have requested multiple files, you can
specify a corrected file name or you can enter "IGNORE" to continue
accessing the remaining file names entered.
=> file pctfull
COST IN U.S. DOLLARS
                                                 SINCE FILE
                                                                 TOTAL
                                                      ENTRY
                                                               SESSION
FULL ESTIMATED COST
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                                                                 48.90
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DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)
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                                                      ENTRY
                                                               SESSION
CA SUBSCRIBER PRICE
                                                       0.00
                                                                 -1.50
FILE 'PCTFULL' ENTERED AT 11:54:09 ON 25 APR 2006
COPYRIGHT (C) 2006 Univentio
FILE LAST UPDATED:
                           25 APR 2006
                                            <20060425/UP>
MOST RECENT UPDATE WEEK:
                               200616
                                             <200616/EW>
FILE COVERS 1978 TO DATE
>>> IMAGES ARE AVAILABLE ONLINE AND FOR EMAIL-PRINTS <<<
>>> NEW IPC8 DATA AND FUNCTIONALITY NOW AVAILABLE IN THIS FILE.
    SEE
    http://www.stn-international.de/stndatabases/details/ipc-reform.html >>>
>>> FOR CHANGES IN PCTFULL PLEASE SEE HELP CHANGE
    (last updated April 10, 2006) <<<
=> s E1-E3
         43187 "PYRIDINE"/BI
       1001528 "2"/BI
        984281 "4"/BI
         16650 "DICARBOXYLIC"/BI
        266611 "ACID"/BI
        204039 "BIS"/BI
         15856 "BIPHENYL"/BI
       1001528 "2"/BI
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8080 "YLMETHYL"/BI 60149 "AMIDE"/BI

)"/BI

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             4 449734/BI
        389660 45/BI
        964506 6/BI
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                 ((449734(W)45(W)6)/BI)
             1 544678/BI
        181577 73/BI
       1009273 1/BI
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L5
             0 ("PYRIDINE-2,4-DICARBOXYLIC ACID BIS(((BIPHENYL)-2-YLMETHYL)AMID
              E) "/BI OR 449734-45-6/BI OR 544678-73-1/BI)
=> file derwent
'DERWENT' IS NOT A VALID FILE NAME
SESSION CONTINUES IN FILE 'PCTFULL'
Enter "HELP FILE NAMES" at an arrow prompt (=>) for a list of files
that are available. If you have requested multiple files, you can
specify a corrected file name or you can enter "IGNORE" to continue
accessing the remaining file names entered.
=> file uspat2 epfull
COST IN U.S. DOLLARS
                                                 SINCE FILE
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                                                      ENTRY
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                                                                 53.54
FULL ESTIMATED COST
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                                                                 TOTAL
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)
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CA SUBSCRIBER PRICE
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FILE 'USPAT2' ENTERED AT 11:56:41 ON 25 APR 2006
CA INDEXING COPYRIGHT (C) 2006 AMERICAN CHEMICAL SOCIETY (ACS)
FILE 'EPFULL' ENTERED AT 11:56:41 ON 25 APR 2006
COPYRIGHT (C) 2006 European Patent Office / FIZ Karlsruhe
=> s E1-E3
             2 ("PYRIDINE-2,4-DICARBOXYLIC ACID BIS(((BIPHENYL)-2-YLMETHYL)AMID
L6
               E) "/BI OR 449734-45-6/BI OR 544678-73-1/BI)
=> d L6 1-2 ti abs bib
     ANSWER 1 OF 2 USPAT2 on STN
       Pyridine matrix metalloproteinase inhibitors
TI
       Selective MMP-13 inhibitors are pyridine derivatives of the formula
AB
                or a pharmaceutically acceptable salt thereof, wherein:
R.sup.1 and R.sup.2 independently are hydrogen, halo, hydroxy, C.sub.1-C.sub.6
       alkyl, C.sub.1-C.sub.6 alkoxy, C.sub.2-C.sub.6 alkenyl, C.sub.2-C.sub.6
       alkynyl, NO.sub.2, NR.sup.4R.sup.5, CN, or CF.sub.3;
E is independently 0 or S;
A and B independently are OR.sup.4 or NR.sup.4R.sup.5;
R.sup.4 and R.sup.5 independently are H, C.sub.1-C.sub.6 alkyl, C.sub.2-C.sub.6
       alkenyl, C.sub.2-C.sub.6 alkynyl, (CH.sub.2).sub.n aryl,
       (CH.sub.2).sub.n cycloalkyl, (CH.sub.2).sub.n heteroaryl, or R.sup.4 and
       R.sup.5 when taken together with the nitrogen to which they are attached
       complete a 3- to 8-membered ring containing carbon atoms and optionally
       containing a heteroatom selected from O, S, or NH, and optionally
       substituted or unsubstituted;
n is an integer of from 0 to 6.
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       2004:268379 USPAT2
AN
       Pyridine matrix metalloproteinase inhibitors
ΤI
TN
       Barvian, Nicole Chantel, Ann Arbor, MI, UNITED STATES
```

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Connor, David Thomas, Ann Arbor, MI, UNITED STATES
       O'Brien, Patrick Michael, Stockbridge, MI, UNITED STATES
       Ortwine, Daniel Fred, Saline, MI, UNITED STATES
       Patt, William Chester, Chelsea, MI, UNITED STATES
       Wilson, Michael William, Ann Arbor, MI, UNITED STATES
PΑ
       Warner-Lambert Company, Morris Plains, NJ, UNITED STATES (U.S.
       corporation)
                          В2
                               20060321
PΤ
      US 7015237
      US 2004-842863
                               20040510 (10)
AΙ
      Division of Ser. No. US 2002-71073, filed on 8 Feb 2002, Pat. No. US
RLI
       6881743
PRAI
      US 2001-268781P
                           20010214 (60)
ידת
      Utility
FS
       GRANTED
EXNAM Primary Examiner: Morris, Patricia L.
       Purchase, Jr., Claude F., Crissey, Todd M., Ashbrook, Charles W.
LREP
CLMN
       Number of Claims: 5
ECL
       Exemplary Claim: 1
DRWN
      No Drawings
LN.CNT 1497
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
     ANSWER 2 OF 2 USPAT2 on STN
       Pyridine matrix metalloproteinase inhibitors
ΤI
       Selective MMP-13 inhibitors are pyridine derivatives of the formula
AB
       ##STR1##
                 or a pharmaceutically acceptable salt thereof,
wherein:
R.sup.1 and R.sup.2 independently are hydrogen, halo, hydroxy, C.sub.1-C.sub.6
       alkyl, C.sub.1-C.sub.6 alkoxy, C.sub.2-C.sub.6 alkenyl, C.sub.2-C.sub.6
       alkynyl, NO.sup.2, NR.sup.4R.sup.5, CN, or CF.sub.3,
E is independently 0 or S;
A and B independently are OR.sup.4 or NR.sup.4R.sup.5;
R.sup.4 and R.sup.5 independently are H, C.sub.1-C.sub.6 alkyl, C.sub.2-C.sub.6
       alkenyl, C.sub.2-C.sub.6 alkynyl, (CH.sub.2).sub.n aryl,
       (CH.sub.2).sub.n cycloalkyl, (CH.sub.2).sub.n heteroaryl, or R.sup.4 and
       R.sup.5 when taken together with the nitrogen to which they are attached
       complete a 3- to 8-membered ring containing carbon atoms and optionally
       containing a heteroatom selected from O, S, or NH, and optionally
       substituted or unsubstituted,
n is an integer of from 0 to 6.
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       2002:288130 USPAT2
       Pyridine matrix metalloproteinase inhibitors
TI
IN
       Barvian, Nicole Chantel, Ann Arbor, MI, UNITED STATES
       Connor, David Thomas, Ann Arbor, MI, UNITED STATES
       O'Brien, Patrick Michael, Stockbridge, MI, UNITED STATES
       Ortwine, Daniel Fred, Saline, MI, UNITED STATES
       Patt, William Chester, Chelsea, MI, UNITED STATES
       Shuler, Kevon Ray, Chelsea, MI, UNITED STATES
       Wilson, Michael William, Ann Arbor, MI, UNITED STATES
PA
       Warner-Lambert Company, Morris Plains, NJ, UNITED STATES (U.S.
       corporation)
                               20050419
PT
       US 6881743
                          B2
       US 2002-71073
                               20020208 (10)
AΙ
                          20010214 (60)
PRAI
       US 2001-268781P
       Utility
DT
FS
       GRANTED
       Primary Examiner: Morris, Patricia L.
EXNAM
       Pfizer Inc., Ashbrook, Charles W., Purchase, Jr., Claude F.
LREP
CLMN
       Number of Claims: 13
ECL
       Exemplary Claim: 1
DRWN
       0 Drawing Figure(s); 0 Drawing Page(s)
LN.CNT 1604
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                                                                SESSION
CA SUBSCRIBER PRICE
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                                                                  -1.50
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                                     <20060416/UP>
FILE LAST UPDATED: 16 APR 2006
PATENTS CITATION INDEX, COVERS 1973 TO DATE
>>> LEARNING FILE LDPCI AVAILABLE <<<
>>> PLEASE BE AWARE OF THE NEW IPC REFORM IN 2006, SEE
http://www.stn-international.de/stndatabases/details/ipc reform.html and
http://scientific.thomson.com/media/scpdf/ipcrdwpi.pdf <<<
=> s E1-E3
          6788 "PYRIDINE"/BI
        141300 "2"/BI
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        183896 "ACID"/BI
         14095 "BIS"/BI
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        141300 "2"/BI
           119 "YLMETHYL"/BI
         12498 "AMIDE"/BI
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                 BIPHENYL" (W) "2" (W) "YLMETHYL" (W) "AMIDE") /BI)
             0 449734/BI
          1482 45/BI
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             0 449734-45-6/BI
                 ((449734(W)45(W)6)/BI)
             0 544678/BI
            58 73/BI
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                 ((544678(W)73(W)1)/BI)
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L7
               E) "/BI OR 449734-45-6/BI OR 544678-73-1/BI)
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FILE 'DRUGMONOG' ACCESS NOT AUTHORIZED
COST IN U.S. DOLLARS
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FULL ESTIMATED COST
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CA SUBSCRIBER PRICE
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FILE 'WPIFV' ENTERED AT 11:58:24 ON 25 APR 2006
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FILE 'WPINDEX' ACCESS NOT AUTHORIZED
=> index E1-E3
'PYRIDINE-2,4' IS NOT A VALID FILE NAME
ENTER A FILE NAME OR (IGNORE):
ENTER A FILE NAME OR (IGNORE):ignore
'544678-73-1' IS NOT A VALID FILE NAME
ENTER A FILE NAME OR (IGNORE):ignore
AT LEAST TWO FILES ARE NEEDED FOR STNINDEX
SESSION CONTINUES IN THE PREVIOUS FILE ENVIRONMENT
To see a list of files and clusters that may be used in the INDEX
command, enter HELP FILE NAMES or HELP CLUSTER NAMES at an arrow
prompt. Only the learning files and NBSFLUIDS may not be used in
STNINDEX.
=> focus E1-E3
'E1-E3' IS NOT VALID HERE
Only numbers and dashes (-) may be used to specify the answers to
be FOCUSed, for example, "FOCUS L2 1-10 13".
ENTER ANSWER NUMBERS OR TAGGED (?):
ENTER ANSWER NUMBERS OR TAGGED (?):
ENTER ANSWER NUMBERS OR TAGGED (?):1
PROCESSING COMPLETED FOR L6
L8
              1 FOCUS L6 1
=> s E1-E3
  14 FILES SEARCHED...
  31 FILES SEARCHED...
  46 FILES SEARCHED...
  64 FILES SEARCHED...
            10 ("PYRIDINE-2,4-DICARBOXYLIC ACID BIS(((BIPHENYL)-2-YLMETHYL)AMID
               E) "/BI OR 449734-45-6/BI OR 544678-73-1/BI)
=> dup rem L9
DUPLICATE IS NOT AVAILABLE IN 'ADISINSIGHT, ADISNEWS, DGENE, DRUGMONOG2,
FOREGE, GENBANK, IMSPRODUCT, IMSRESEARCH, KOSMET, NUTRACEUT, PCTGEN, PHAR,
PHARMAML, PROUSDDR, PS, RDISCLOSURE, SYNTHLINE'.
ANSWERS FROM THESE FILES WILL BE CONSIDERED UNIQUE
PROCESSING COMPLETED FOR L9
              6 DUP REM L9 (4 DUPLICATES REMOVED)
=> d L10 1-6 ti abs bib
    ANSWER 1 OF 6 USPATFULL on STN
                                                         DUPLICATE 1
TΤ
       Pyridine matrix metalloproteinase inhibitors
       Selective MMP-13 inhibitors are pyridine derivatives of the formula
AB
       ##STR1##
       or a pharmaceutically acceptable salt thereof,
       wherein:
       R.sup.1 and R.sup.2 independently are hydrogen, halo, hydroxy,
       C.sub.1-C.sub.6 alkyl, C.sub.1-C.sub.6 alkoxy, C.sub.2-C.sub.6 alkenyl,
       C.sub.2-C.sub.6 alkynyl, N0.sub.2, NR.sup.4R.sup.5, CN, or CF.sub.3;
```

E is independently O or S;

A and B independently are OR.sup.4 or NR.sup.4R.sup.5;

R.sup.4 and R.sup.5 independently are H, C.sub.1-C.sub.6 alkyl, C.sub.2-C.sub.6 alkenyl, C.sub.2-C.sub.6 alkynyl, (CH.sub.2).sub.n aryl, (CH.sub.2).sub.n cycloalkyl, (CH.sub.2).sub.n heteroaryl, or R.sup.4 and R.sup.5 when taken together with the nitrogen to which they are attached complete a 3- to 8-membered ring containing carbon atoms and optionally containing a heteroatom selected from O, S, or NH, and optionally substituted or unsubstituted;

n is an integer of from 0 to 6.

```
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
      2004:268379 USPATFULL
      Pyridine matrix metalloproteinase inhibitors
TI
IN
      Barvian, Nicole Chantel, Ann Arbor, MI, UNITED STATES
      Connor, David Thomas, Ann Arbor, MI, UNITED STATES
      O'Brien, Patrick Michael, Stockbridge, MI, UNITED STATES
      Ortwine, Daniel Fred, Saline, MI, UNITED STATES
      Patt, William Chester, Chelsea, MI, UNITED STATES
      Wilson, Michael William, Ann Arbor, MI, UNITED STATES
PΙ
      US 2004209922
                          A1
                               20041021
      US 7015237
                          B2
                               20060321
ΑI
      US 2004-842863
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                               20040510 (10)
RLI
      Division of Ser. No. US 2002-71073, filed on 8 Feb 2002, PENDING
PRAI
      US 2001-268781P
                           20010214 (60)
DT
      Utility
FS
      APPLICATION
      WARNER-LAMBERT COMPANY, 2800 PLYMOUTH RD, ANN ARBOR, MI, 48105
LREP
CLMN
      Number of Claims: 13
ECL
      Exemplary Claim: CLM-001-9
DRWN
      No Drawings
LN.CNT 1657
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
```

- L10 ANSWER 2 OF 6 USPATFULL on STN
- TI Combination of an allosteric carboxylic inhibitor of matrix metalloproteinase-13 with a selective inhibitor of cyclooxygenase-2 that is not celecoxib or valdecoxib
- This invention provides a combination, comprising an allosteric AB carboxylic inhibitor of MMP-13, or a pharmaceutically acceptable salt thereof, with a selective inhibitor of COX-2, or a pharmaceutically acceptable salt thereof, that is not celecoxib or valdecoxib. This invention also provides a method of treating a disease that is responsive to inhibition of MMP-13 and cyclooxygenase-2, comprising administering to a patient suffering from such a disease the invention combination comprising an allosteric carboxylic inhibitor of MMP-13, or a pharmaceutically acceptable salt thereof, with a selective inhibitor of COX-2, or a pharmaceutically acceptable salt thereof, that is not celecoxib or valdecoxib. This invention also provides a pharmaceutical composition, comprising the invention combination comprising an allosteric carboxylic inhibitor of MMP-13, or a pharmaceutically acceptable salt thereof, with a selective inhibitor of COX-2, or a pharmaceutically acceptable salt thereof, that is not celecoxib or valdecoxib, and a pharmaceutically acceptable carrier, diluent, or excipient. This invention also provides a combination comprising an NSAID, or a pharmaceutically acceptable salt thereof, and an allosteric carboxylic inhibitor of MMP-13, or a pharmaceutically acceptable salt thereof. This invention also provides a pharmaceutical composition, comprising the invention combination comprising an allosteric carboxylic inhibitor of MMP-13, or a pharmaceutically acceptable salt thereof, with an NSAID, or a pharmaceutically acceptable salt thereof, and a

pharmaceutically acceptable carrier, diluent, or excipient. This invention also provides a method of treating a disease that is responsive to inhibition of MMP-13 and cyclooxygenase-1 or cyclooxygenase-2, comprising administering to a patient suffering from such a disease the invention combination comprising an allosteric carboxylic inhibitor of MMP-13, or a pharmaceutically acceptable salt thereof, with an NSAID, or a pharmaceutically acceptable salt thereof. The invention combinations may also be further combined with other pharmaceutical agents depending on the disease being treated.

CAS INDEXING IS AVAILABLE FOR THIS PATENT. 2004:25212 USPATFULL TΙ Combination of an allosteric carboxylic inhibitor of matrix metalloproteinase-13 with a selective inhibitor of cyclooxygenase-2 that is not celecoxib or valdecoxib Roark, William Howard, Ann Arbor, MI, UNITED STATES IN PΙ US 2004019054 A1 20040129 ΑI US 2003-619769 A1 20030715 (10) PRAI US 2002-396785P 20020717 (60) DT Utility FS APPLICATION WARNER-LAMBERT COMPANY, 2800 PLYMOUTH RD, ANN ARBOR, MI, 48105 LREP CLMN Number of Claims: 9 ECL Exemplary Claim: 1 DRWN No Drawings LN.CNT 8368 CAS INDEXING IS AVAILABLE FOR THIS PATENT. ANSWER 3 OF 6 USPATFULL on STN ΤI Combination of an allosteric carboxylic inhibitor of matrix metalloproteinase-13 with celecoxib or valdecoxib This invention provides a combination, comprising an allosteric ΔR carboxylic inhibitor of MMP-13, or a pharmaceutically acceptable salt thereof, with celecoxib, or a pharmaceutically acceptable salt thereof, or valdecoxib, or a pharmaceutically acceptable salt thereof. This invention also provides a method of treating a disease that is responsive to inhibition of MMP-13 and cyclooxygenase-2, comprising administering to a patient suffering from such a disease the invention combination comprising an allosteric carboxylic inhibitor of MMP-13, or a pharmaceutically acceptable salt thereof, with celecoxib, or a pharmaceutically acceptable salt thereof, or valdecoxib, or a pharmaceutically acceptable salt thereof. This invention also provides a pharmaceutical composition, comprising the invention combination comprising an allosteric carboxylic inhibitor of MMP-13, or a pharmaceutically acceptable salt thereof, with celecoxib, or a pharmaceutically acceptable salt thereof, or valdecoxib, or a

The invention combination may also be further combined with other pharmaceutical agents depending on the disease being treated.

pharmaceutically acceptable salt thereof, and a pharmaceutically

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CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       2004:25211 USPATFULL
AΝ
ΤI
       Combination of an allosteric carboxylic inhibitor of matrix
      metalloproteinase-13 with celecoxib or valdecoxib
IN
      Roark, William Howard, Ann Arbor, MI, UNITED STATES
PΙ
      US 2004019053
                         A1
                              20040129
AΙ
      US 2003-619662
                        A1
                               20030715 (10)
      US 2002-396903P
                         20020717 (60)
PRAI
DT
      Utility
FS
      APPLICATION
LREP
      WARNER-LAMBERT COMPANY, 2800 PLYMOUTH RD, ANN ARBOR, MI, 48105
CLMN
      Number of Claims: 9
ECL
      Exemplary Claim: 1
```

acceptable carrier, diluent, or excipient.

DRWN No Drawings LN.CNT 8040 CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L10 ANSWER 4 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN

TI Preparation of 2,4-pyridinedicarboxamides and 4,6-pyrimidinedicarboxamides as inhibitors of collagenase (MMP 13)

GI

$$R^1$$
 R^2
 R^3
 R^2
 R^3
 R^2

AB Title compds. [I; A = CH, N; R1-R3 = H, halo, (halogenated) alkyl, alkoxy, OH, CO2R4, cyano, NR5R6, etc.; R4 = H, alkyl; R5, R6 = H, alkyl, alkylcarbonyl, etc.; or R1R2, R2R3 = 5-6 membered (aromatic) (saturated) (hetero)cyclyl], were prepd for the treatment of degenerative joint diseases. Thus, 4,6-pyrimidinedicarboxylic acid in SOCl2 was stirred for 2 h at 85° followed by addition of CH2Cl2 at room temperature and Et3N at 0°. The reaction mixture was further stirred with 3-chloro-4-fluorobenzylamine for 15 min to give 40% N,N-bis(3-chloro-4-fluorobenzyl)pyrimidine-4,6-dicarboxamide. The latter inhibited collagenase 3 (MMP 13) with IC50 = 23 nM.

I

- AN 2003:467290 CAPLUS
- DN 139:53028
- TI Preparation of 2,4-pyridinedicarboxamides and 4,6-pyrimidinedicarboxamides as inhibitors of collagenase (MMP 13)
- IN Habermann, Joerg; Weithmann, Klaus-Ulrich; Kogler, Herbert; Kirsch, Reinhard; Wehner, Volkmar
- PA Aventis Pharma Deutschland G.m.b.H., Germany
- SO Ger. Offen., 20 pp.

CODEN: GWXXBX

DT Patent

LA German

FAN.CNT 1

	PA:	ATENT NO. 				APPLICATION NO.						DATE						
PI		1016 2469															0011	
	WO	2003	0497	38		A1		2003	0619	1	WO 2	002-	EP13:	240		20	0021	125
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	EΡ	1455	790			A1		2004	0915		EP 2	002-	7927	99		20	0021	125
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	JP	2005	5130	49		T2		2005	0512		JP 2	003-	5507	87		20	0021	125
	US	2003	2291							1	US 2	002-	6599	4		20	0021	209
	US	6933	298			B2		2005	0823									
PRAI		2001																
	US	2002	-358	887P		P		2002	0222									

OS MARPAT 139:53028

L10 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 2
TI Preparation of pyridinedicarboxamide and -dicarboxylic acid derivatives as selective MMP-13 matrix metalloproteinase inhibitors with therapeutic uses

GI

$$\begin{array}{c|c}
R^2 \\
R^1 \\
C \\
C \\
R \\
E \\
E
\end{array}$$

Selective MMP-13 inhibitors are pyridine derivs. (I; e.g. AB pyridine-2,4-dicarboxylic acid bis(3-methoxybenzylamide)) or a pharmaceutically acceptable salt thereof, wherein: R1 and R2 independently are H, halo, hydroxy, C1-C6 alkyl, C1-C6 alkoxy, C2-C6 alkenyl, C2-C6 alkynyl, NO2, NR4R5, CN, or CF3; E is independently O or S; A and B independently are OR4 or NR4R5; R4 and R5 independently are H, C1-C6 alkyl, C2-C6 alkenyl, C2-C6 alkynyl, (CH2)n aryl, (CH2)n cycloalkyl, (CH2)n heteroaryl, or R4 and R5 when taken together with the N to which they are attached complete a 3- to 8-membered ring containing C atoms and optionally containing a heteroatom selected from O, S, or NH, and optionally substituted or unsubstituted; n is 0 to 6. Although I and other Markush structures in the patent show 2,4- derivs., many specific 3,5- derivs. are included in the claims and examples. Combinatorial and non-combinatorial methods were used to prepare numerous claimed compds. and characterization data is reported for about 90 compds. IC50 values for various claimed compds. show the selectivity towards MMP-13 vs. MMP-1 and MMP-3 and the potent MMP-13 inhibitory activity (e.g. 0.033 µM for pyridine-2,4-dicarboxylic acid bis[((1,3-benzodioxol-5-yl)methyl)amide]).

AN 2002:637657 CAPLUS

DN 137:185420

TI Preparation of pyridinedicarboxamide and -dicarboxylic acid derivatives as selective MMP-13 matrix metalloproteinase inhibitors with therapeutic uses

IN Barvian, Nicole Chantel; Connor, David Thomas; O'brien, Patrick Michael; Ortwine, Daniel Fred; Patt, William Chester; Shuler, Kevon Ray; Wilson, Michael William

PA Warner-Lambert Company, USA

SO PCT Int. Appl., 68 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

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	EP 1362033							EP 2002-716263						20020204			

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PRAI US 2001-268781P
                        P
                              20010214
                        W
    WO 2002-IB345
                              20020204
    US 2002-71073
                        A3
                              20020208
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MARPAT 137:185420

RE.CNT 10 THERE ARE 10 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 6 OF 6 IFIPAT COPYRIGHT 2006 IFI on STN DUPLICATE 3 L10

PYRIDINE MATRIX METALLOPROTEINASE INHIBITORS; TREATING DISEASES RESULTING ΤI FROM TISSUE BREAKDOWN SUCH AS HEART DISEASE, MULTIPLE SCLEROSIS, OSTEO-AND RHEUMATOID ARTHRITIS, ATHEROSCLEROSIS, AND OSTEOPOROSIS

Selective MMP-13 inhibitors are pyridine derivatives of the formula AΒ

DRAWING

or a pharmaceutically acceptable salt thereof, wherein: R1 and R2 independently are hydrogen, halo, hydroxy, C1-C6 alkyl, C1-C6 alkoxy, C2-C6 alkenyl, C2-C6 alkynyl, NO2, NR4R5, CN, or CF3, E is independently O or S; A and B independently are OR4 or NR4R5; R4 and R5 independently are H. C1-C6 alkyl, C2-C6 alkenyl, C2-C6 alkynyl, (CH2)n aryl, (CH2)n cycloalkyl, (CH2)n heteroaryl, or R4 and R5 when taken together with the nitrogen to which they are attached complete a 3 to 8-membered ring containing carbon atoms and optionally containing a heteroatom selected from O, S, or NH, and optionally substituted or unsubstituted, n is an integer of from 0 to 6.

CLMN

AN 10217293 IFIPAT; IFIUDB; IFICDB

PYRIDINE MATRIX METALLOPROTEINASE INHIBITORS; TREATING DISEASES RESULTING ΤI FROM TISSUE BREAKDOWN SUCH AS HEART DISEASE, MULTIPLE SCLEROSIS, OSTEO-AND RHEUMATOID ARTHRITIS, ATHEROSCLEROSIS, AND OSTEOPOROSIS

INF Barvian; Nicole Chantel, Ann Arbor, MI, US Connor; David Thomas, Ann Arbor, MI, US O'Brien; Patrick Michael, Stockbridge, MI, US Ortwine; Daniel Fred, Saline, MI, US Patt; William Chester, Chelsea, MI, US Shuler; Kevon Ray, Chelsea, MI, US Wilson; Michael William, Ann Arbor, MI, US

IN Barvian Nicole Chantel; Connor David Thomas; O'Brien Patrick Michael; Ortwine Daniel Fred; Patt William Chester; Shuler Kevon Ray; Wilson Michael William

Unassigned PAF

Unassigned Or Assigned To Individual (68000)

PPA Warner-Lambert Co (Probable)

AG Claude F. Purchase, Jr. Warner-Lambert Company, 2800 Plymouth Road, Ann Arbor, MI 48105, US

PΙ US 2002161000 A1 20021031 ΑI US 2002-71073 20020208

PRAI US 2001-268781P 20010214 (Provisional)

US 2002161000 20021031 US 6881743 20050419

DTUtility; Patent Application - First Publication

FS CHEMICAL

APPLICATION

CLMN 35

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ALL L# QUERIES AND ANSWER SETS ARE DELETED AT LOGOFF

LOGOFF? (Y)/N/HOLD:y

COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 178.83 241.26 FULL ESTIMATED COST

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)

SINCE FILE TOTAL
ENTRY SESSION
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NEWS 3 DEC 23 New IPC8 SEARCH, DISPLAY, and SELECT fields in USPATFULL/
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NEWS
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NEWS 5 JAN 13
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                 INPADOC
NEWS
     6 JAN 17
                 Pre-1988 INPI data added to MARPAT
     7 JAN 17 IPC 8 in the WPI family of databases including WPIFV
NEWS
NEWS
     8 JAN 30 Saved answer limit increased
     9 FEB 21 STN AnaVist, Version 1.1, lets you share your STN AnaVist
NEWS
                 visualization results
NEWS 10 FEB 22
                 The IPC thesaurus added to additional patent databases on STN
NEWS 11 FEB 22
                 Updates in EPFULL; IPC 8 enhancements added
NEWS 12 FEB 27
                 New STN AnaVist pricing effective March 1, 2006
NEWS 13 FEB 28 MEDLINE/LMEDLINE reload improves functionality
NEWS 14 FEB 28 TOXCENTER reloaded with enhancements
NEWS 15 FEB 28 REGISTRY/ZREGISTRY enhanced with more experimental spectral
                 property data
NEWS 16 MAR 01 INSPEC reloaded and enhanced
NEWS 17 MAR 03 Updates in PATDPA; addition of IPC 8 data without attributes
NEWS 18 MAR 08 X.25 communication option no longer available after June 2006
NEWS 19 MAR 22 EMBASE is now updated on a daily basis
NEWS 20 APR 03 New IPC 8 fields and IPC thesaurus added to PATDPAFULL
NEWS 21 APR 03 Bibliographic data updates resume; new IPC 8 fields and IPC
                 thesaurus added in PCTFULL
NEWS 22 APR 04
                 STN AnaVist $500 visualization usage credit offered
NEWS 23 APR 12
                 LINSPEC, learning database for INSPEC, reloaded and enhanced
                 Improved structure highlighting in FQHIT and QHIT display
NEWS 24 APR 12
                 in MARPAT
                 Derwent World Patents Index to be reloaded and enhanced during
NEWS 25
         APR 12
                 second quarter; strategies may be affected
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NEWS EXPRESS FEBRUARY 15 CURRENT VERSION FOR WINDOWS IS V8.01a,
CURRENT MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP),
AND CURRENT DISCOVER FILE IS DATED 19 DECEMBER 2005.
V8.0 AND V8.01 USERS CAN OBTAIN THE UPGRADE TO V8.01a AT
http://download.cas.org/express/v8.0-Discover/

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COST IN U.S. DOLLARS

COST IN U.S. DOLLARS
SINCE FILE TOTAL
ENTRY SESSION
FULL ESTIMATED COST
0.21
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FILE 'CAPLUS' ENTERED AT 15:59:56 ON 25 APR 2006 .
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=> s pyridinedicarbox? op pyridinecarboxy?

5068 PYRIDINEDICARBOX?

14670 OP

5743 PYRIDINECARBOXY?

L1 0 PYRIDINEDICARBOX? OP PYRIDINECARBOXY? (PYRIDINEDICARBOX? (W) OP (W) PYRIDINECARBOXY?)

=> s pyridinedicarbox? or pyridinecarboxy?

5068 PYRIDINEDICARBOX?

5743 PYRIDINECARBOXY?

L2 10247 PYRIDINEDICARBOX? OR PYRIDINECARBOXY?

=> s L2 and (solubilizing(w)agent)

10403 SOLUBILIZING

775866 AGENT

1385 SOLUBILIZING (W) AGENT

L3 3 L2 AND (SOLUBILIZING (W) AGENT)

=> d L3 1-3 ti abs bib

L3 ANSWER 1 OF 3 CAPLUS COPYRIGHT 2006 ACS on STN

TI Pharmaceutical compositions containing polymer for enhanced drug concentrations

AB A drug in a solubility-improved form is combined with a concentration-enhancing polymer, i.e., a cellulosic or non-cellulosic polymer, in a sufficient amount so that the combination provides substantially enhanced drug concentration

in a use environment,, such as digestive tract, s.c. space, vagina, lung, blood vessels, and muscle relative to a control comprising the same amount

of the same solubility-improved form of drug without the concentration-enhancing

polymer. For example, the solubility of sertraline-HCl was increased in presence of citric acid, giving a solubility-improvement factor of 9.3. Thus, citric acid is an excellent solubilizing agent for sertraline-HCl. A solution was prepared containing 1000 μg/mL sertraline-HCl, 500 μg/mL citric acid, and 1000 μg/mL hydroxypropyl Me cellulose acetate succinate (HPMCAS) in phosphate buffer. (pH 7.9). Addition of the concentration-enhancing polymer HPMCAS resulted in a maximum concentration that was

1.7-fold that of control containing no polymer.

AN 2001:489208 CAPLUS

DN 135:97443

TI Pharmaceutical compositions containing polymer for enhanced drug concentrations

IN Babcock, Walter Christian; Curatolo, William John; Friesen, Dwayne Thomas; Lorenz, Douglas Alan; Nightingale, James Alan Schriver; Shanker, Ravi Mysore

PA Pfizer Products Inc., USA

SO PCT Int. Appl., 85 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

				D DATE	APPLICATION NO.				
ΡI					WO 2000-IB1787				
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		CR, CU,	CZ, DE,	DK, DM, DZ,	EE, ES, FI, GB, GD, GE,	GH, GM, HR,			
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	BR 2000	016555	Α	20020917	BR 2000-16555	20001201			
	EP 1239	835	A1	20020918	EP 2000-976217	20001201			
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		01617			TR 2002-200201617	20001201			
		518485			JP 2001-548090	20001201			
	EE 2002	00357	Α	20031015	EE 2002-357	20001201			
	AU 7843		B2		AU 2001-14091	20001201			
	US 2002	006443			US 2000-742785	20001220			
	BG 1067	64	Α	20030331	BG 2002-106764	20020531			
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PRAI	US 1999	-171841P	P	19991223					
	WO 2000	-IB1787	W	20001201					

RE.CNT 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L3 ANSWER 2 OF 3 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Chelation extraction of lead from soil using pyridine-2,6-dicarboxylic
- AB Pyridine-2,6-dicarboxylic acid (PDA) was used as a Pb-complexing agent.

 Batch testing involving PDA extraction of Pb from spiked soils showed PDA to be an effective Pb solubilizing agent across a wide pH

range. Pb extraction efficiency was independent of the total carbonate concentration,

competing cations, or the soil aging period. PDA compared favorably with EDTA as a Pb-complexing agent, while behaving more desirably than EDTA in releasing the extracted Pb. PDA was effectively reclaimed and reused in 4

successive extraction procedures, achieving in each run a Pb extraction efficiency that exceeded 80% recovery of the total Pb present in the spiked soil. In recovery procedures, the complex solution was elevated to a pH of .apprx.10, separating the Pb as a hydroxide precipitate, and allowing for virtually complete recovery of the PDA in solution ΑN 1995:414081 CAPLUS DN 122:195965 Chelation extraction of lead from soil using pyridine-2,6-dicarboxylic TI Macauley, Edward; Hong, Andrew AU Department of Civil Engineering, University of Utah, Salt Lake City, UT, CS 84112, USA Journal of Hazardous Materials (1995), 40(3), 257-70 SO CODEN: JHMAD9; ISSN: 0304-3894 PB Elsevier DTJournal LΑ English ANSWER 3 OF 3 CAPLUS COPYRIGHT 2006 ACS on STN L3 Solubilizing agents. VIII. Pharmaceutical application of acid amides as ΤI solubilizing agents AB cf. CA 55, 10440c. Ten acid amides were examined for their use as a solubilization agent. The majority of them have very small toxicity. Solubilization effects of these substances on sparingly soluble pharmaceutical prepns. and food additives showed that they can be used for such practical purposes. 1962:13216 CAPLUS DN 56:13216 OREF 56:2515e-f Solubilizing agents. VIII. Pharmaceutical application of acid amides as solubilizing agents ΑU Samejima, Masayoshi CS Tanabe Siyaku Co., Osaka SO Yakugaku Zasshi (1961), 81, 1208 CODEN: YKKZAJ; ISSN: 0031-6903 DTJournal LA Unavailable => s pyridinedicarboxamide or pyridinecarboxamide 309 PYRIDINEDICARBOXAMIDE 1547 PYRIDINECARBOXAMIDE 1836 PYRIDINEDICARBOXAMIDE OR PYRIDINECARBOXAMIDE L4=> s L4 and (solubilizing(w)agent) 10403 SOLUBILIZING 775866 AGENT 1385 SOLUBILIZING (W) AGENT L5 1 L4 AND (SOLUBILIZING(W) AGENT) => d 15 ti abs bib ANSWER 1 OF 1 CAPLUS COPYRIGHT 2006 ACS on STN L5 Solubilizing agents. VIII. Pharmaceutical application of acid amides as ΤI solubilizing agents AB cf. CA 55, 10440c. Ten acid amides were examined for their use as a solubilization agent. The majority of them have very small toxicity. Solubilization effects of these substances on sparingly soluble pharmaceutical prepns. and food additives showed that they can be used for such practical purposes.

AN

DN

1962:13216 CAPLUS

56:13216 OREF 56:2515e-f

- TI Solubilizing agents. VIII. Pharmaceutical application of acid amides as solubilizing agents
- AU Samejima, Masayoshi
- CS Tanabe Siyaku Co., Osaka
- SO Yakugaku Zasshi (1961), 81, 1208 CODEN: YKKZAJ; ISSN: 0031-6903
- DT Journal
- LA Unavailable
- => s L4 and arthritis
 - 40833 ARTHRITIS
- L6 42 L4 AND ARTHRITIS
- => d L6 1-46 ti
- L6 ANSWER 1 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Adamantyl derivatives as P2X7 receptor antagonists, their preparation, pharmaceutical compositions, and use in therapy
- L6 ANSWER 2 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Pyridine derivatives, particularly 6-amino-5-phenylpyridine-2-carboxylic acid amides, with activity as sodium channel modulators, useful for the treatment of pain, and their preparation, pharmaceutical compositions, and use
- L6 ANSWER 3 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Preparation of nitrogen-heteroaryl-containing protein kinase modulators for use against cancer and other diseases
- L6 ANSWER 4 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Preparation of triazolyl arylbenzamides as inhibitors of cytokines
- L6 ANSWER 5 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Preparation of pyridine derivatives as akt kinase inhibitors
- L6 ANSWER 6 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Preparation of 2-pyridinecarboxamides as kinase inhibitors
- L6 ANSWER 7 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Preparation of amido-substituted indazoles as Rho-kinase inhibitors
- L6 ANSWER 8 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Preparation of benzyl ether amine compounds useful as CCR-5 antagonists
- L6 ANSWER 9 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Preparation of aminofurazanyl imidazopyridines as Rho kinase inhibitors
- L6 ANSWER 10 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Fluoro substituted omega-carboxyaryl diphenyl urea for treatment of raf, VEGFR, PDGFR, p38 and flt-3 kinase-mediated diseases
- L6 ANSWER 11 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Preparation of pyrimidinecarboxamides, pyrimidinylcarbamates and related compounds as inhibitors of T cell activation for the treatment of inflammatory diseases
- L6 ANSWER 12 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI A preparation of (thiopyranyloxy)pyridinecarboxamide derivatives, useful as $TNF-\alpha$ and PDE4 inhibitors
- L6 ANSWER 13 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI A preparation of amphiphilic pyridinium compounds, useful for suppression of IL-8 secretion

- L6 ANSWER 14 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Preparation of imidazo-fused oxazolo[4,5-b]pyridine and imidazo-fused thiazolo[4,5-b]pyridine based tricyclic compounds as IKK kinase inhibitors for treating inflammation and immune disorders
- L6 ANSWER 15 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI A pharmaceutical composition comprising adamantane derivative P2X7 antagonists and sulfasalazine
- L6 ANSWER 16 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Preparation of 2-anilino-4-(imidazol-5-yl)pyrimidine derivatives and their use as cdk (cdk2) kinase inhibitors
- L6 ANSWER 17 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Preparation of bicyclic (hetero)aryl- and pyridine-containing diaryl ureas as Raf kinase and angiogenesis inhibitors useful in the treatment of cancer and other disorders
- L6 ANSWER 18 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Preparation of 2-oxo-1,3,5-perhydrotriazapine derivatives for treatment of hyper-proliferative, angiogenesis, and inflammatory disorders
- L6 ANSWER 19 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Preparation of compounds having 4-pyridylalkylthio group as inhibitors of angiogenesis and vascular permeability
- L6 ANSWER 20 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Preparation of indazole derivatives as JNK enzyme inhibitors
- L6 ANSWER 21 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Preparation of (pyridyl)(phenylpyridyl)pyrazoles as inhibitors of the transforming growth factor β
- L6 ANSWER 22 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Preparation of 2-phenylpyridin-4-yl heterocycles as selective activin-like kinase-5 inhibitors useful against fibrosis and other disorders
- L6 ANSWER 23 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Preparation of aminothiazoles as inhibitors of the transforming growth factor-beta (TGF- β) signalling pathway
- L6 ANSWER 24 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Preparation of aryl ureas with angiogenesis inhibiting activity
- L6 ANSWER 25 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Preparation of 2,4-pyridinedicarboxamides and 4,6-pyrimidinedicarboxamides as inhibitors of collagenase (MMP 13)
- L6 ANSWER 26 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Preparation of isoindoles as Factor Xa inhibitors
- L6 ANSWER 27 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Preparation of 3-(azahetero)aryl-1H-pyrazolo[3,4-d]pyrimidin-3-amines as protein kinase inhibitors with antiangiogenic properties
- L6 ANSWER 28 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Preparation of 3-(azahetero)aryl-1H-pyrazolo[3,4-d]pyrimidin-3-amines as protein kinase inhibitors with antiangiogenic properties
- L6 ANSWER 29 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Preparation of 3-(azahetero)aryl-1H-pyrazolo[3,4-d]pyrimidin-3-amines as protein kinase inhibitors with antiangiogenic properties
- L6 ANSWER 30 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Preparation of benzimidazol derivatives as modulators of chemokine

receptors

- L6 ANSWER 31 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Preparation of **pyridinedicarboxamide** and -dicarboxylic acid derivatives as selective MMP-13 matrix metalloproteinase inhibitors with therapeutic uses
- L6 ANSWER 32 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Preparation of carbamoyl-substituted pyridinyl aryl ether derivatives as inhibitors of phosphodiesterase IV isozymes
- L6 ANSWER 33 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Preparation of novel N-substituted- γ , γ -trisubstituted lactam derivatives as matrix metalloproteinase inhibitors
- L6 ANSWER 34 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors
- L6 ANSWER 35 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Preparation of cyclic amine derivatives for inhibition of the action of chemokines such as MIP-1 α and/or MCP-1 on target cells
- L6 ANSWER 36 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Preparation of Furanoisoquinoline derivatives as phosphodiesterase IV inhibitors
- L6 ANSWER 37 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Use of arylalkanoylpyridazines for treatment of osteoporosis, tumors, arteriosclerosis, rheumatoid arthritis, multiple sclerosis, diabetes mellitus, ulcerative colitis and AIDS
- L6 ANSWER 38 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Preparation of heterocyclic glycyl β -alanine derivatives as vitronectin antagonists
- L6 ANSWER 39 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI 1,2-Substituted imidazolyl compounds for the treatment of inflammation
- L6 ANSWER 40 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Dietary supplement for pain relief
- L6 ANSWER 41 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Heteroaryl-Fused 2-Phenylisothiazolone Inhibitors of Cartilage Breakdown
- L6 ANSWER 42 OF 42 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Transformations in the pyridine series. A simple preparation of 3-methyl-4-phenylpyridine and corresponding 2-carboxamides
- => s L6 anf py<2002 MISSING OPERATOR L6 ANF The search profile that was entered contains terms or nested terms that are not separated by a logical operator.
- => s L6 and py<2002 21808456 PY<2002
- L7 7 L6 AND PY<2002
- => d L7 1-7 ti abs bib
- L7 ANSWER 1 OF 7 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Preparation of Furanoisoquinoline derivatives as phosphodiesterase IV inhibitors

AB Title compds. [I; R1 = C6H5, 4-HOC6H4, 1-naphthyl, 4-CH3OC6H4, 2-CH3OC6H4, 4-NH2C6H4, 4-C6H5C6H4, 4-BrC6H4, CH3, C6H5CO, 3-CH3SCH2CONHC6H4, 3-CH3OCOC6H4, 3-NH2C(CH3)2CONHC6H4, 3-furyl, 3-HOOCC6H4, 2-chloro-4-pyridyl, 3-CH3CH2OCOC6H4, 4-pyridylethylaminocarbonyl; R2 = CH3, CH2Br, CH3CH2, H, CH3COO; R3 = CH3, H; R2R3 = (CH2)5; R4 = H, CH2N(CH3)2, CH2SC6H5, CH2C(:CH2)CH3, CH2NHCOCH3, CH3OCH2, CH2OH, CH2F, CH2COOH, CH2CN; R5 = C1, OCH3, CON(CH3)2, CH3O, H, CH3CH2O, NH2, CHONH, CH3SO2NH, NH2CONH, CH3CH2S, CH3; R6 = CH3, H, CH3CH; R7 = CH3, H, CH3CH2; R6R7 = (CH2)5; R8 = H, CH3; R9 = H, CH3; Y = CH2, CHOH, C:O, C(CH3)2; X =electron pair, 0, S; n = 0, 1] and salts are prepared as phosphodiesterase IV inhibitors. Title compds. are useful as preventives and remedies for diseases caused by inflammation, for example, bronchial asthma, chronic obstructive pulmonary disease (COPD), rheumatoid arthritis, autoimmune disease and diabetes. Thus, the title compound I (R6 = CH3; R7 = CH3; R2 = CH3; R3 = CH3; X = O; R5 = CH3; n = O; R9 = H; R8 = H; R1 = CH3; R1 = CH3; R2 = CH3; R3 = CH3; R4 = CH3; R5 = CH33-CH3S:OCH2CONHC6H4) was prepared and biol. tested.

ΑN 2001:713354 CAPLUS

DN 135:272895

- Preparation of Furanoisoquinoline derivatives as phosphodiesterase IV TI inhibitors
- Kawano, Yasuhiko; Matsumoto, Tatsumi; Uchikawa, Osamu; Fujii, Nobuhiro; IN Tarui, Naoki
- Takeda Chemical Industries, Ltd., USA PA
- SO PCT Int. Appl., 620 pp.

CODEN: PIXXD2

DTPatent

LA Japanese

FAN.CNT 1

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PATENT NO.
                         KIND
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                                            AU 2001-39550
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                                20030102
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PRAI JP 2000-87121
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     WO 2001-JP2277
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     CASREACT 135:272895; MARPAT 135:272895
RE.CNT 11
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THERE ARE 11 CITED REFERENCES AVAILABLE FOR THIS RECORD

ALL CITATIONS AVAILABLE IN THE RE FORMAT

L7 ANSWER 2 OF 7 CAPLUS COPYRIGHT 2006 ACS on STN

TI Use of arylalkanoylpyridazines for treatment of osteoporosis, tumors, arteriosclerosis, rheumatoid arthritis, multiple sclerosis, diabetes mellitus, ulcerative colitis and AIDS

GI

$$R^{2}$$
 $N-N$
 Q
 $N+COB$
 T

The use of 1-acyl-3-aryl-1,4,5,6-tetrahydropyridazines I (B = A, OA, NH2, NHA, NHAA'; A,A' = C1-10-alkyl, fluoro-C1-10-alkyl, chloro-C1-10-alkyl, (substituted) heterocyclyl; Q = absent, C1-6-alkylene; R1, R2 = OH, OR5, SR5, SOR5, SO2R5, halo, NO2, NH2, NHR5, NR5R6, or R1R2 = OCH2O; R5, R6 = alkyl, C3-7-cycloalkyl, C4-8-methylenecycloalkyl, C2-8-alkenylene) is claimed for producing a medicament for treating osteoporosis, tumors, arteriosclerosis, rheumatoid arthritis, multiple sclerosis, diabetes mellitus, ulcerative colitis and AIDS. Thus, 4-[(nicotinoyl)amino]benzoyl chloride was added to a mixture of potassium tert-butylate and 3-(3,4-dimethoxyphenyl)-1,4,5,6-tetrahydropyridazine in THF to give N-[4-[[3-(3,4-dimethoxyphenyl)-5,6-dihydro-1(4H)-pyridazinyl]carbonyl]phenyl]-3-pyridinecarboxamide hydrochloride. The pharmacol. activity of the claimed compds. was not shown.

AN 2000:725445 CAPLUS

DN 133:301117

TI Use of arylalkanoylpyridazines for treatment of osteoporosis, tumors, arteriosclerosis, rheumatoid arthritis, multiple sclerosis, diabetes mellitus, ulcerative colitis and AIDS

IN Rochus, Jonas; Wolf, Michael; Beier, Norbert; Kluxen, Franz-Werner;
Fittschen, Claus

PA Merck Patent G.m.b.H., Germany

SO PCT Int. Appl., 34 pp. CODEN: PIXXD2

DT Patent

LA German

FAN.CNT 1

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	PATENT NO.					KINI)]	DATE		1	APPL:	ICAT:	ION 1	NO.		DATE			
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	EP	1143	944			A2	:	2001	1017]	EP 20	000-	9169	49		20	0000	315 <-	-
	EP	1143	944			A3	:	2002	0911										
		R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT,	

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IE, SI, LT, LV, FI, RO
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     JP 2002541095
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     NO 2001004845
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PRAI DE 1999-19915364
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     WO 2000-EP2280
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     MARPAT 133:301117
OS
     ANSWER 3 OF 7 CAPLUS COPYRIGHT 2006 ACS on STN
L7
     Preparation of heterocyclic glycyl \beta-alanine derivatives as
TI
     vitronectin antagonists
AΒ
     Tile compds. A(CY3Z3)t-Het-CO-V-(CYZ)n-CONR11CHR1(CH2)pCOR [Het =
     (un) substituted 5-8 membered monocyclic heterocyclic ring containing 1-4
     heteroatoms selected from O, N, or S, optionally unsatd. and linked to
     (CY3Z3)t and CO at the 1- and 3-positions; A = NR5C(:Y1)NR7R8,
     NR5C(:NR7)Y2, or N:C(NR2R5)(NR7R8), where Y1 = NR2, O, S; R2, R7, R8 = H,
     alkyl, aryl, amino, etc. or R2 and R8 taken together form an
     (un) substituted dinitrogen heterocycle; R5 = H, alkyl, alkenyl, alkynyl,
     benzyl, phenethyl; and Y2 = alkyl, cycloalkyl, bicycloalkyl, aryl, etc.; V
     = NR6, where R6 = H, alkyl, cycloalkyl, aralkyl, aryl, monocyclic
     heterocyclyl or R6 together with Y forms a mono-nitrogen-containing ring; Y,
     Y3, Z, Z3 = H, alkyl, aryl, cycloalkyl or Y and Z together or Y3 and Z3
     together form cycloalkyl; n = 1-3; t = 0-2; p = 0-3; R = X-R3, where X =
     O, S, or NR4 and R3 and R4 = H, alkyl, sugars, steroids, etc.; R1 = H,
     alkyl, alkenyl, alkynyl, aryl, etc.] or their pharmaceutically acceptable
     salts were prepared as vitronectin antagonists. Thus, 5-
     [(aminoiminomethyl)amino]-N-[2-[[2-carboxy-1-(3-bromo-5-chloro-2-
     hydroxyphenyl) ethyl] amino] -2-oxoethyl] -3-pyridinecarboxamide
     bis(trifluoroacetate) was prepared and showed IC50 = 1.58 nM for inhibition
     of human vitronectin receptor (\alpha v\beta 3).
     1999:672798 CAPLUS
AN
DN
     131:299691
     Preparation of heterocyclic glycyl β-alanine derivatives as
TI
     vitronectin antagonists
     Chandrakumar, Nizal Samuel; Desai, Bipinchandra Nanubhai; Devadas,
IN
     Balekudru; Huff, Renee; Khanna, Ish K.; Rao, Shashidhar N.; Rico, Joseph
     G.; Rogers, Thomas E.; Ruminski, Peter G.; Russell, Mark Andrew; Yu, Yi;
     Gasiecki, Alan Frank; Malecha, James W.; Miyashiro, Julie M.
PA
     G.D. Searle and Co., USA
SO
     PCT Int. Appl., 269 pp.
     CODEN: PIXXD2
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FAN.CNT 1
     PATENT NO.
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             MD, RU, TJ, TM
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JP 2000-543454

19990409

JP 2002511462

F	RU 2215746	C2	20031110	RU	2000-128033	19990409	
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ľ	NO 2000005084	Α	20001127	NO	2000-5084	20001009	<
τ	US 2004127477	A1	20040701	US	2003-718328	20031120	
PRAI (US 1998-81394P	P	19980410				
Ţ	US 1999-289140	A3	19990408				
V	WO 1999-US4297	W	19990409				
os N	MARPAT 131:299691						

RE.CNT 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

L7 ANSWER 4 OF 7 CAPLUS COPYRIGHT 2006 ACS on STN
TI 1,2-Substituted imidazolyl compounds for the treatment of inflammation
GI

$$F_3C$$
 N
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 SO_2R^7
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 SO_2Me
 II

A class of imidazolyl compds., which are selective inhibitors of AB cyclooxygenase 2 (COX 2), is described. The compds. are useful in treating inflammation and related disorders (arthritis, fever, and pain). Compds. of particular interest are I [R3 = H, (un)substituted alkyl, aralkyl, heterocycloalkyl, acyl, cyano, alkoxy, alkylthio, cycloalkoxy, halo, substituted carbonyl, sulfonyl, oxy, thio, aryl, and heteroaryl; R7 = alkyl or amino; R8 = ≥ 1 of H, halo, alkyl, haloalkyl, alkoxy, amino, haloalkoxy, cyano, CO2H, OH, hydroxyalkyl, alkoxyalkyl, alkylamino, nitro, and alkylthio], as well as certain heterocyclic analogs. For instance, condensation of 4-(methylsulfonyl)aniline-HCl with 3-cyanopyridine in the presence of Me3Al (34%), followed by cyclization of the resultant amidine with BrCH2COCF3 (60%), and dehydration of the obtained hydroxydihydroimidazole derivative using p-MeC6H4SO3H (23%), gave title compound II. In the carrageenan-induced rat paw edema and analgesia tests, II gave 57% inhibition of edema at 30 mg/kg orally, and 51% inhibition of hyperalgesic foot withdrawal at 10 mg/kg orally. Inhibition data for recombinant COX 1 and 2 are also given.

AN 1996:363276 CAPLUS

DN 125:33646

TI 1,2-Substituted imidazolyl compounds for the treatment of inflammation

IN Khanna, Ish K.; Weier, Richard M.; Collins, Paul W.; Yu, Yi; Xu, Xiangdong; Huff, Renee M.; Partis, Richard A.; Koszyk, Francis J.

PA G.D. Searle and Co., USA

SO PCT Int. Appl., 249 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 3

	PAT	ENT	NO.			KIN	D	DATE		i	APPL	ICAT:	ION 1	. 07		D	ATE		
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PI	WO	9603	388			A1		1996	0208	1	WO 1	995-1	US95	06		1	9950'	727	<
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     JP 10503211
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    AT 224374
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                          B2
                                            AU 2001-11100
                                                                    20010109
    AU 767993
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    AU 1997-15739
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    WO 1997-US300
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                                19970124
                          B1
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    US 1999-101493
    US 2001-4944
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                                20011205
                                20030902
    US 2003-653399
                          A1
os
    MARPAT 125:33646
     ANSWER 5 OF 7 CAPLUS COPYRIGHT 2006 ACS on STN
L7
TI
    Dietary supplement for pain relief
     A dietary supplement containing sources of vitamins B3, B5, and/or B6,
AΒ
     D-phenylalanine, glucosamine sulfate, and optionally mucopolysaccharides
     such as chondroitin sulfate and shark cartilage can provide relief of
     joint or muscular pain, e.g. arthritis. Thus, a tablet
     formulation contained pantothenic acid 100, shark cartilage 100,
     DL-phenylalanine 50, chondroitin sulfate 50, glucosamine sulfate 50 mg,
     and conventional tableting additives.
     1995:958356 CAPLUS
AN
DN
     123:350289
     Dietary supplement for pain relief
ΤI
IN
     Woodward, Robert John
PA
SO
     Brit. UK Pat. Appl., 9 pp.
```

CODEN: BAXXDU

- DTPatent
- English LΑ
- FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
ΡI	GB 2286528	A1	19950823	GB 1994-3063	19940217 <
	GB 2286528	B2	19980916		
PRAI	GB 1994-3063		19940217		

- ANSWER 6 OF 7 CAPLUS COPYRIGHT 2006 ACS on STN L7
- Heteroaryl-Fused 2-Phenylisothiazolone Inhibitors of Cartilage Breakdown ΤI GI

AB The synthesis, biol. evaluation, and structure-activity relationships of a series of N-Ph heteroaryl-fused isothiazolones, e.g. I (R = aryl) are described. These isothiazolones have been shown to exhibit potent, dose-dependent inhibition of IL-1 β -induced breakdown of proteoglycan in a cartilage organ culture assay. This effect is likely due to inhibition of MMP activation and a consequent reduction in MMP activity following IL-1 β stimulation. Thus these compds. potentially represent simple, non-peptidic disease-modifying agents for the treatment of arthritic diseases. To examine the effects of structure on in vitro activity, three general features of the mols. were varied, substituents on the pendant N-Ph group, the position of ring fusion to the isothiazolone, and substituents on the fused ring peri to the isothiazolone sulfur.

AN 1994:605254 CAPLUS

DN 121:205254

TI Heteroaryl-Fused 2-Phenylisothiazolone Inhibitors of Cartilage Breakdown AU Wright, Stephen W.; Petraitis, Joseph J.; Abelman, Matthew M.; Batt,

Douglas G.; Bostrom, Lori L.; Corbett, Ronald L.; Decicco, Carl P.; Di Meo, Susan V.; Freimark, Bruce; et al.

Inflammatory Diseases Research, DuPont Merck Pharmaceutical Company,

II

Wilmington, DE, 19880-0353, USA Journal of Medicinal Chemistry (**1994**), 37(19), 3071-8

CODEN: JMCMAR; ISSN: 0022-2623

DT Journal

CS

SO

GI

LA English

L7 ANSWER 7 OF 7 CAPLUS COPYRIGHT 2006 ACS on STN

TI Transformations in the pyridine series. A simple preparation of

3-methyl-4-phenylpyridine and corresponding 2-carboxamides

AB 3-Methyl-4-phenylpyridine (I) was prepared in 96% yield by Huang-Minlon reduction of 4-phenyl-3-pyridinecarboxaldehyde. I was carboxylated followed by amidation to give its 1-pyridinecarboxamides, e.g. II. At 30 mg/kg II reduced edema in the rat adjuvant arthritis test by 17%.

AN 1980:550093 CAPLUS

DN 93:150093

TI Transformations in the pyridine series. A simple preparation of 3-methyl-4-phenylpyridine and corresponding 2-carboxamides

AU Harrison, Ernest A., Jr.; Rice, Kenner C.; Rogers, Michael E.

CS Natl. Inst. Arthritis Metab. Dig. Dis., NIH, Bethesda, MD, 20205, USA

SO Heterocycles (1980), 14(6), 813-16 CODEN: HTCYAM; ISSN: 0385-5414

DT Journal

LA English

=> s pyridinedicarboxamide

L8 309 PYRIDINEDICARBOXAMIDE

=> s L8 and inflamm?

241376 INFLAMM?

L9 5 L8 AND INFLAMM?

=> s L8 and ?nflamm?

260115 ?NFLAMM?

L10 5 L8 AND ?NFLAMM?

=> s L10 and py<2003 22795154 PY<2003

L11 5 L10 AND PY<2003

=> d L11 1-5 ti abs bib

L11 ANSWER 1 OF 5 CAPLUS COPYRIGHT 2006 ACS on STN

TI Preparation of **pyridinedicarboxamide** and -dicarboxylic acid derivatives as selective MMP-13 matrix metalloproteinase inhibitors with therapeutic uses

GΙ

$$\begin{array}{c|c}
R^2 \\
R^1 \\
C \\
C \\
R \\
E \\
E
\end{array}$$

Selective MMP-13 inhibitors are pyridine derivs. (I; e.g. AB pyridine-2,4-dicarboxylic acid bis(3-methoxybenzylamide)) or a pharmaceutically acceptable salt thereof, wherein: R1 and R2 independently are H, halo, hydroxy, C1-C6 alkyl, C1-C6 alkoxy, C2-C6 alkenyl, C2-C6 alkynyl, NO2, NR4R5, CN, or CF3; E is independently O or S; A and B independently are OR4 or NR4R5; R4 and R5 independently are H, C1-C6 alkyl, C2-C6 alkenyl, C2-C6 alkynyl, (CH2)n aryl, (CH2)n cycloalkyl, (CH2)n heteroaryl, or R4 and R5 when taken together with the N to which they are attached complete a 3- to 8-membered ring containing C atoms and optionally containing a heteroatom selected from O, S, or NH, and optionally substituted or unsubstituted; n is 0 to 6. Although I and other Markush structures in the patent show 2,4- derivs., many specific 3,5- derivs. are included in the claims and examples. Combinatorial and non-combinatorial methods were used to prepare numerous claimed compds. and characterization data is reported for about 90 compds. IC50 values for various claimed compds. show the selectivity towards MMP-13 vs. MMP-1 and MMP-3 and the potent MMP-13 inhibitory activity (e.g. 0.033 µM for pyridine-2,4-dicarboxylic acid bis[((1,3-benzodioxol-5-yl)methyl)amide]).

AN 2002:637657 CAPLUS

DN 137:185420

TI Preparation of **pyridinedicarboxamide** and -dicarboxylic acid derivatives as selective MMP-13 matrix metalloproteinase inhibitors with therapeutic uses

IN Barvian, Nicole Chantel; Connor, David Thomas; O'brien, Patrick Michael; Ortwine, Daniel Fred; Patt, William Chester; Shuler, Kevon Ray; Wilson, Michael William

PA Warner-Lambert Company, USA

SO PCT Int. Appl., 68 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

	PATENT NO.					KIND DATE			APPLICATION NO.					DATE					
Ρľ	WO 2002064568				A1 20020822			WO 2002-IB345					20020204 <						
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    EE 200300391
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    MARPAT 137:185420
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RE.CNT 10 THERE ARE 10 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L11 ANSWER 2 OF 5 CAPLUS COPYRIGHT 2006 ACS on STN
- TI New N-aryl-2,3-pyridinedicarboxamides: chemical and pharmacological studies
- AB Six title compds. were prepared by dehydration of 2,3-pyridinedicartboxylic acid with acetic anhydride and subsequent reactions with aromatic amines to yield amides. The products were characterized by spectrometric methods and their toxicity and analgesic and anti-inflammatory effects were tested in male mice. Antibacterial and antifungal activities were tested in vitro.
- AN 1992:440006 CAPLUS
- DN 117:40006
- TI New N-aryl-2,3-pyridinedicarboxamides: chemical and pharmacological studies
- AU Albuquerque, C. N.; Bacha, C. T. M.; Schapoval, E. E. S.; Loiseau, P.; Flores, S. B.
- CS Fac. Farmacia, UFRGS, Porto Alegre, Brazil
- SO Revista Brasileira de Farmacia (1991), 72(2), 31-3 CODEN: RBFAAH; ISSN: 0370-372X
- DT Journal
- LA Portuguese
- L11 ANSWER 3 OF 5 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Pyridinedicarboxamides

GI

AB Title compds. I (R = halo, NHNH2, amino, OR1, SR1; R1 = alkyl, cycloalkyl, alkenyl, alkynyl, aralkyl, Ph, pyridyl), useful as inflammation

inhibitors, analgesics, antithrombotics, and neoplasm inhibitors (no data), were prepared Thus, treating 8 g II with MeOH and 40% MeNH2 gave 7.1 g I (R = Cl).

AN 1984:51459 CAPLUS

DN 100:51459

TI Pyridinedicarboxamides

PA Chugai Pharmaceutical Co., Ltd., Japan

SO Jpn. Kokai Tokkyo Koho, 5 pp.

CODEN: JKXXAF

DT Patent

LA Japanese

FAN.CNT 1

I'AN.	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
ΡI	JP 58159465	A2	19830921	JP 1982-40902	19820317 <
	JP 01042267	B4	19890911		
PRAT	JP 1982-40902		19820317		

L11 ANSWER 4 OF 5 CAPLUS COPYRIGHT 2006 ACS on STN

TI Pyridinedicarboxamides

GI

Two pyridinedicarboxamides I [R = 2-HO2CC6H4 (II), 2-pyridyl] were prepared by condensation of 2,6-pyridinedicarbonyl dichloride (III) with 2-H2NC6H4CO2H (IV) or 2-aminopyridine. I had antiinflammatory, antipyretic, and analgesic activities (no data). Thus, 2.04 g III was treated with 2.7 g IV and 1 g NaOH at 20° to give 1.9 g II.

AN 1977:468163 CAPLUS

DN 87:68163

TI Pyridinedicarboxamides

IN Matsuzaki, Meiki; Okabe, Hiroshi; Tanaka, Seishiro

PA Banyu Pharmaceutical Co., Ltd., Japan

SO Jpn. Kokai Tokkyo Koho, 2 pp.

CODEN: JKXXAF

DT Patent

LA Japanese

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	JP 52033676	A2	19770314	JP 1975-108495	19750909 <
PRAI	JP 1975-108495	Α	19750909		•

- L11 ANSWER 5 OF 5 CAPLUS COPYRIGHT 2006 ACS on STN
- TI Antiinflammatory p-aminosalicylic acid derivatives
- GI For diagram(s), see printed CA Issue.

AB Amines I (R1 = Ph, pyridyl, thienyl, pyrrolyl, furyl; R2 = H, Me, Et, Bu, Pr, iso-Pr, Na, K, Ca, Mg, Al) were prepared by acylating 4-aminosalicyclic acid (II) or its esters with acids R1CO2H or acid chlorides R1COCl. I were effective against carrageenin-induced edema in rats (ED30 20-120 mg/kg) and had low toxicity in mice (LD50 >3 g/kg). Thus, 3.8 g II in aqueous NaOH was acylated with 3.04 g 2,6-pyridinedicarbonyl chloride to give 71.1% III (R2 = H), converted to di-Na salt. Also prepared was III (R2 = Et).

AN 1975:409805 CAPLUS

DN 83:9805

TI Antiinflammatory p-aminosalicylic acid derivatives

IN Matsuzaki, Meiki; Okabe, Hiroshi; Tanaka, Seishiro; Nakamura, Koichi;

Yoshida, Akio PA Banyu Pharmaceutical Co., Ltd. SO Jpn. Kokai Tokkyo Koho, 4 pp. CODEN: JKXXAF DT Patent LA Japanese FAN.CNT 1 KIND APPLICATION NO. DATE PATENT NO. DATE ---------------JP 49110641 A2 19741022 JP 1973-28009 19730312 <--JP 57015584 B4 19820331 PRAI JP 1973-28009 A 19730312 => s pyridinedicarboxamide and (antibiotic or antimicrobial or antiviral) 309 PYRIDINEDICARBOXAMIDE 124384 ANTIBIOTIC 62249 ANTIMICROBIAL 53225 ANTIVIRAL L12 2 PYRIDINEDICARBOXAMIDE AND (ANTIBIOTIC OR ANTIMICROBIAL OR ANTIVI RAL) => d L12 1-2 ti abs bib L12 ANSWER 1 OF 2 CAPLUS COPYRIGHT 2006 ACS on STN New N-aryl-2,3-pyridinedicarboxamides: chemical and pharmacological studies Six title compds. were prepared by dehydration of 2,3-pyridinedicartboxylic AB acid with acetic anhydride and subsequent reactions with aromatic amines to yield amides. The products were characterized by spectrometric methods and their toxicity and analgesic and anti-inflammatory effects were tested in male mice. Antibacterial and antifungal activities were tested in vitro. 1992:440006 CAPLUS AN DN117:40006 New N-aryl-2,3-pyridinedicarboxamides: chemical and pharmacological TI studies ΑU Albuquerque, C. N.; Bacha, C. T. M.; Schapoval, E. E. S.; Loiseau, P.; Flores, S. B. CS Fac. Farmacia, UFRGS, Porto Alegre, Brazil Revista Brasileira de Farmacia (1991), 72(2), 31-3 SO CODEN: RBFAAH; ISSN: 0370-372X DTJournal LA Portuguese L12 ANSWER 2 OF 2 CAPLUS COPYRIGHT 2006 ACS on STN Kanamycin derivatives TI Kanamycin (I) derivs. which maintain high blood levels of the AB antibiotic for prolonged periods of time are prepared by treating I with at least 4 moles of a monocyclic aromatic monoaldehyde. Thus, 11.4 g. I in 50 ml. H2O added to 7 g. p-ClC6H4CHO in 50 ml. MeOH caused the temperature to rise to 40-50° and an oil to deposit. The oil crystallized on scratching. The product, after standing overnight, was filtered off, treated with C in hot MeOH and the MeOH was diluted with H2O until crystallization was initiated. The crystalline product was dried 2 hrs. at 61° in vacuo to yield the I tetrakis(p-chlorobenzylidene) derivative (II) (containing 3.88% H2O), m. 213-16° (decomposition). Several analogs of II were prepared similarly (starting aldehyde and m.p. of analog given): o-HOC6H4CHO, above 165° (decomposition); p-HOC6H4CHO, 193-6° (decomposition); p-Me2NC6H4CHO, 255-8°; 3,4-(MeO)2C6H3CHO, 173-5° (decomposition); BzH, 235-7° (decomposition); p-MeC6H4CHO, -; p-O2NC6H4CHO, -; 2-pyridinecarboxaldehyde, -. Derivs. of furfural and 2-

thiophenecarboxaldehyde were also used.

1961:2773 CAPLUS

AN

DN 55:2773

OREF 55:574g-i

TI Kanamycin derivatives

PA Bristol Laboratories International, S.A.

DT Patent

LA Unavailable

FAN.CNT 1

PATENT NO. KIND DATE APPLICATION NO. DATE

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COST IN U.S. DOLLARS

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FULL ESTIMATED COST

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)

SINCE FILE TOTAL
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SINCE FILE
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SESSION
CA SUBSCRIBER PRICE

-13.50

FILE 'MEDLINE' ENTERED AT 16:07:22 ON 25 APR 2006

FILE LAST UPDATED: 22 APR 2006 (20060422/UP). FILE COVERS 1950 TO DATE.

On December 11, 2005, the 2006 MeSH terms were loaded.

The MEDLINE reload for 2006 is now (26 Feb.) available. For details on the 2006 reload, enter HELP RLOAD at an arrow prompt (=>). See also:

http://www.nlm.nih.gov/mesh/

http://www.nlm.nih.gov/pubs/techbull/nd04/nd04_mesh.html

http://www.nlm.nih.gov/pubs/techbull/nd05/nd05_med_data_changes.html

http://www.nlm.nih.gov/pubs/techbull/nd05/nd05_2006_MeSH.html

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MEDLINE thesauri in the /CN, /CT, and /MN fields incorporate the MeSH 2006 vocabulary.

This file contains CAS Registry Numbers for easy and accurate substance identification.

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COST IN U.S. DOLLARS

SINCE FILE TOTAL
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FULL ESTIMATED COST

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CA SUBSCRIBER PRICE

SINCE FILE TOTAL
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STRUCTURE FILE UPDATES: 24 APR 2006 HIGHEST RN 881733-90-0 DICTIONARY FILE UPDATES: 24 APR 2006 HIGHEST RN 881733-90-0

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TSCA INFORMATION NOW CURRENT THROUGH January 6, 2006

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* The CA roles and document type information have been removed from *
* the IDE default display format and the ED field has been added, *
* effective March 20, 2005. A new display format, IDERL, is now *
* available and contains the CA role and document type information. *

Structure search iteration limits have been increased. See HELP SLIMITS for details.

REGISTRY includes numerically searchable data for experimental and predicted properties as well as tags indicating availability of experimental property data in the original document. For information on property searching in REGISTRY, refer to:

http://www.cas.org/ONLINE/UG/regprops.html

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E1 THROUGH E6 ASSIGNED

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http://www.nlm.nih.gov/mesh/

http://www.nlm.nih.gov/pubs/techbull/nd04/nd04 mesh.html

http://www.nlm.nih.gov/pubs/techbull/nd05/nd05 med data changes.html

http://www.nlm.nih.gov/pubs/techbull/nd05/nd05_2006_MeSH.html

OLDMEDLINE is covered back to 1950.

MEDLINE thesauri in the /CN, /CT, and /MN fields incorporate the MeSH 2006 vocabulary.

This file contains CAS Registry Numbers for easy and accurate substance identification.

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            0 VALECOXIB/BI
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       122706 IMMUNODEFICIENCY
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L15 ANSWER 1 OF 4
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     Gateways to clinical trials.
     Gateways to Clinical Trials is a guide to the most recent clinical trials
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     in current literature and congresses. The data in the following tables
     have been retrieved from the Clinical Trials Knowledge Area of Prous
     Science Integrity, the drug discovery and development portal,
     http://integrity.prous.com. This issue focuses on the following selection
     of drugs: Abiraterone acetate, acyline, adalimumab, adenosine
     triphosphate, AEE-788, AIDSVAX gp120 B/B, AK-602, alefacept, alemtuzumab,
     alendronic acid sodium salt, alicaforsen sodium, alprazolam, amdoxovir,
     AMG-162, aminolevulinic acid hydrochloride, aminolevulinic acid methyl
     ester, aminophylline hydrate, anakinra, anecortave acetate, anti-CTLA-4
     MAb, APC-8015, aripiprazole, aspirin, atazanavir sulfate, atomoxetine
     hydrochloride, atorvastatin calcium, atrasentan, AVE-5883, AZD-2171;
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     cisplatin, clofarabine, clopidogrel hydrogensulfate, clozapine,
     co-trimoxazole, CP-122721, creatine, CY-2301, cyclophosphamide, cypher,
     cytarabine, cytolin; D0401, darbepoetin alfa, darifenacin hydrobromide,
     DASB, desipramine hydrochloride, desloratadine, desvenlafaxine succinate,
     dexamethasone, didanosine, diquafosol tetrasodium, docetaxel, doxorubicin
     hydrochloride, drotrecogin alfa (activated), duloxetine hydrochloride,
     dutasteride; Ecallantide, efalizumab, efavirenz, eletriptan,
     emtricitabine, enfuvirtide, enoxaparin sodium, estramustine phosphate
     sodium, etanercept, ethinylestradiol, etonogestrel,
     etonogestrel/ethinylestradiol, etoposide, exenatide; Famciclovir,
     fampridine, febuxostat, filgrastim, fludarabine phosphate, fluocinolone
     acetonide, fluorouracil, fluticasone propionate, fluvastatin sodium,
     fondaparinux sodium; Gaboxadol, gamma-hydroxybutyrate sodium, gefitinib,
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gelclair, gemcitabine, gemfibrozil, glibenclamide, glyminox; Haloperidol, heparin sodium, HPV 16/HPV 18 vaccine, human insulin, human insulin; Icatibant, imatinib mesylate, indium 111 (111In) ibritumomab tiuxetan, infliximab, INKP-100, iodine (I131) tositumomab, IoGen, ipratropium bromide, ixabepilone; L-870810, lamivudine, lapatinib, laquinimod, latanoprost, levonorgestrel, licochalcone a, liposomal doxorubicin, lopinavir, lopinavir/ritonavir, lorazepam, lovastatin; Maraviroc, maribavir, matuzumab, MDL-100907, melphalan, methotrexate, methylprednisolone, mitomycin, mitoxantrone hydrochloride, MK-0431, MN-001, MRKAd5 HIV-1 gag/pol/nef, MRKAd5gag, MVA.HIVA, MVA-BN Nef, MVA-Muc1-IL-2, mycophenolate mofetil; Nelfinavir mesilate, nesiritide, NSC-330507; Olanzapine, olmesartan medoxomil, omalizumab, oral insulin, osanetant; PA-457, paclitaxel, paroxetine, paroxetine hydrochloride, PCK-3145, PEG-filgrastim, peginterferon alfa-2a, peginterferon alfa-2b, perillyl alcohol, pexelizumab, pimecrolimus, pitavastatin calcium, porfiromycin, prasterone, prasugrel, pravastatin sodium, prednisone, pregabalin, prinomastat, PRO-2000, propofol, prostate cancer vaccine; Rasagiline mesilate, rhBMP-2/ACS, rhBMP-2/BCP, rhC1, ribavirin, rilpivirine, ritonavir, rituximab, Ro-26-9228, rosuvastatin calcium, rosuvastatin sodium, rubitecan; Selodenoson, simvastatin, sirolimus, sitaxsentan sodium, sorafenib, SS(dsFv)-PE38, St. John's Wort extract, stavudine; Tacrolimus, tadalafil, tafenoquine succinate, talaglumetad, tanomastat, taxus, tegaserod maleate, telithromycin, tempol, tenofovir, tenofovir disoproxil fumarate, testosterone enanthate, TH-9507, thalidomide, tigecycline, timolol maleate, tiotropium bromide, tipifarnib, torcetrapib, trabectedin, travoprost, travoprost/timolol, treprostinil sodium; Valdecoxib, vardenafil hydrochloride hydrate, varenicline, VEGF-2 gene therapy, venlafaxine hydrochloride, vildagliptin, vincristine sulfate, voriconazole, VRX-496, VX-385; Warfarin sodium; Ximelagatran; Yttrium 90 (90Y) ibritumomab tiuxetan; Zanolimumab, zidovudine.

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- AN 2005417782 MEDLINE
- DN PubMed ID: 16082422
- TI Gateways to clinical trials.
- AU Bayes M; Rabasseda X; Prous J R
- CS Department of Pharmacology, Prous Science, Barcelona, Spain.. mbayes@prous.com
- SO Methods and findings in experimental and clinical pharmacology, (2005 Jun) Vol. 27, No. 5, pp. 331-72.

 Journal code: 7909595. ISSN: 0379-0355.
- CY Spain
- DT Bibliography
- LA English
- FS Priority Journals
- EM 200509
- ED Entered STN: 6 Aug 2005 Last Updated on STN: 28 Sep 2005 Entered Medline: 27 Sep 2005
- L15 ANSWER 2 OF 4 MEDLINE on STN
- TI Gateways to clinical trials.
- AB Gateways to Clinical Trials is a guide to the most recent clinical trials in current literature and congresses. The data in the following tables has been retrieved from the Clinical Trials Knowledge Area of Prous Science Integrity, the drug discovery and development portal, http://integrity.prous.com. This issue focuses on the following selection of drugs: ABX-IL-8, Acclaim, adalimumab, AGI-1067, alagebrium chloride, alemtuzumab, Alequel, Androgel, anti-IL-12 MAb, AOD-9604, aripiprazole, atomoxetine hydrochloride; Biphasic insulin aspart, bosentan, botulinum toxin type B, bovine lactoferrin, brivudine; Cantuzumab mertansine, CB-1954, CDB-4124, CEA-TRICOM, choriogonadotropin alfa, cilansetron, CpG-10101, CpG-7909, CTL-102, CTL-102/CB-1954; DAC:GRF, darbepoetin alfa, davanat-1, decitabine, del-1 Genemedicine, dexanabinol, dextofisopam, dnaJP1, dronedarone hydrochloride, dutasteride;

Ecogramostim, eletriptan, emtricitabine, EPI-hNE-4, eplerenone, eplivanserin fumarate, erlotinib hydrochloride, ertapenem sodium, escitalopram oxalate, esomeprazole magnesium, etoricoxib, ezetimibe; Falecalcitriol, fingolimod hydrochloride; Gepirone hydrochloride; HBV-ISS, HSV-2 theracine, human insulin; Imatinib mesylate, Indiplon, insulin glargine, ISAtx-247; L612 HuMAb, levodopa/carbidopa/entacapone, lidocaine/prilocaine, LL-2113AD, lucinactant, LY-156735; Meclinertant, metelimumab, morphine hydrochloride, morphine-6-glucuronide; Natalizumab, nimotuzumab, NX-1207, NYVAC-HIV C; Omalizumab, onercept, osanetant; PABA, palosuran sulfate, parathyroid hormone (human recombinant), parecoxib sodium, PBI-1402, PCK-3145, peginterferon alfa-2a, peginterferon alfa-2b, peginterferon alfa-2b/ribavirin, pemetrexed disodium, pimecrolimus, PINC, pregabalin; Ramelteon, rasagiline mesilate, rasburicase, rimonabant hydrochloride, RO-0098557, rofecoxib, rosiglitazone maleate/metformin hydrochloride; Safinamide mesilate, SHL-749, sitaxsentan sodium, sparfosic acid, SprayGel, squalamine, St. John's Wort extract, synthetic human secretin; Taxus, telavancin hydrochloride, telithromycin, temoporfin, tenofovir disoproxil fumarate, tenofovir disoproxil fumarate/emtricitabine, teriparatide, testosterone gel, TG-1024, tirapazamine, travoprost, travoprost/timolol; Valdecoxib, valganciclovir hydrochloride, voriconazole; Ximelagatran.

- AN2005200517 MEDLINE
- PubMed ID: 15834452 DN
- TIGateways to clinical trials.
- ΑU Bayes M; Rabasseda X; Prous J R
- Methods and findings in experimental and clinical pharmacology, (2005 Apr) SO Vol. 27, No. 3, pp. 193-219. Journal code: 7909595. ISSN: 0379-0355.
- CY Spain
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- EM200507
- ED Entered STN: 19 Apr 2005 Last Updated on STN: 16 Jul 2005 Entered Medline: 15 Jul 2005
- ANSWER 3 OF 4 MEDLINE on STN L15
- TI Gateways to clinical trials.
- Gateways to Clinical Trials is a guide to the most recent clinical trials AB reported in current literature and congresses. The data in the following tables have been retrieved from the Clinical Trials Knowledge Area of Prous Science Integrity, the drug discovery and development portal, http://integrity.prous.com. This issue focuses on the following selection of drugs: [188Re]-HDD; A-179578, adalimumab, AK-602, albumin interferon alfa, alfimeprase, amelubant, anakinra, anti-CD2 MAb, APD-356, aripiprazole, atvogen; Bimatoprost, bimosiamose, BLP-25, brivaracetam; Caspofungin acetate, cilansetron, CMV vaccine (bivalent), conivaptan hydrochloride, Cypher; Darbepoetin alfa, darifenacin hydrobromide, D-D4FC, decitabine, dnaJP1, doranidazole, dronedarone hydrochloride; Efalizumab, efaproxiral sodium, emtricitabine, Endeavor, entecavir, erlotinib hydrochloride, escitalopram oxalate, etoricoxib, etravirine, ezetimibe; Fampridine, fenretinide, ferumoxtran-10, forodesine hydrochloride; Gantacurium chloride, gemi-floxacin mesilate, Glyminox, GW-501516; HBV-ISS, hepavir B, human insulin, HuMax-CD20, hyaluronic acid, HyCAMP; Icatibant, IDEA-070, IGN-311, imatinib mesylate, insulin detemir, insulin glargine, insulin qlulisine; Lapatinib, lasofoxifene tartrate, LB-80380, liarozole fumarate, liposome encapsulated doxorubicin, lumiracoxib, LY-570310; MC-1, melatonin, merimepodib, metanicotine, midostaurin; Natalizumab, nicotine conjugate vaccine, NYVAC-HIV C; Patupilone, peginterferon alfa-2a, peginterferon alfa-2b, peginterferon alfa-2b/ribavirin, pelitinib, Peru-15, pexelizumab, PHP, pimecrolimus, prednisolone sodium metasulfobenzoate; Recombinant alfal-antitrypsin (AAT), retigabine, rHA influenza vaccine, rifalazil, rofecoxib,

rosiglitazone maleate/Metformin hydrochloride, rostaporfin, rosuvastatin calcium, rubitecan; Selenite sodium, semilente insulin, SMP-797, sorafenib; Talampanel, tenofovir disoproxil fumarate, TER-199, tiotropium bromide, torcetrapib, treprostinil sodium, TTA; ValboroPro, valdecoxib, val-mCyd, valtorcitabine dihydrochloride: XP-828L. Copyright (c) 2005 Prous Science. All rights reserved.

AN 2005200505 MEDLINE

DN PubMed ID: 15834459

TI Gateways to clinical trials.

AU Bayes M; Rabasseda X; Prous J R

SO Methods and findings in experimental and clinical pharmacology, (2005 Jan-Feb) Vol. 27, No. 1, pp. 49-77.

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ED Entered STN: 19 Apr 2005 Last Updated on STN: 22 Jun 2005 Entered Medline: 21 Jun 2005

- L15 ANSWER 4 OF 4 MEDLINE on STN
- TI Gateways to clinical trials.
- Gateways to Clinical Trials is a guide to the most recent clinical trials AΒ in current literature and congresses. The data in the following tables has been retrieved from the Clinical Studies Knowledge Area of Prous Science Integrity, the drug discovery and development portal, http://integrity.prous.com. This issue focuses on the following selection of drugs: 166Ho-DOTMP 5A8; A-179578, abetimus sodium, adefovir dipivoxil, AGI-1067, AIDSVAX gp120 B/B, AK-602, alefacept alemtuzumab, aliskiren fumarate, ALVAC vCP1433, ALVAC vCP1452, anecortave acetate, arzoxifene hydrochloride, atazanavir sulfate, atlizumab, avasimibe; Binodenoson, BMS-488043; Choriogonadotropin alfa, ciclesonide, COL-1621, CVT-3146, CVT-E002, Cypher; Daptomycin, darbepoetin alfa, darunavir, D-D4FC, deferasirox, desloratadine, desmoteplase, duloxetine hydrochloride, DX-9065a; E-5564, efalizumab, emfilermin, emivirine, emtricitabine, enfuvirtide, estradiol acetate, ezetimibe; Frovatriptan; Gallium maltolate, gefitinib; HIV-1 Immunogen, human insulin; Iquratimod, IL-4/IL-13 Trap, imatinib mesylate, inhaled insulin, insulin qlargine, irofulven, ISS-1018, ivabradine hydrochloride; Lutropin alfa; Melatonin; Nesiritide; O6-Benzylguanine, omapatrilat, oritavancin, ospemifene; Parecoxib sodium, peginterferon alfa-2a, pexelizumab, pimecrolimus, pirfenidone, pramlintide acetate, prasterone sulfate PT-141; Rasburicase, razaxaban hydrochloride, recombinant malaria vaccine, rhBMP-2/ACS, roflumilast, rosiglitazone maleate/metformin hydrochloride, rotavirus vaccine; SCH-D, sitaxsentan sodium, solifenacin succinate; Targinine hydrochloride, taxus, TER-199, tramadol hydrochloride/acetaminophen; Valdecoxib, valganciclovir hydrochloride, vatalanib succinate, VEG Trap(R1R2); Ximelagatran; Yttrium Y90 Epratuzumab.
- AN 2004414825 MEDLINE
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- TI Gateways to clinical trials.
- AU Bayes M; Rabasseda X; Prous J R
- CS Prous Science, S.A., Barcelona, Spain. mbayes@prous.com
- SO Methods and findings in experimental and clinical pharmacology, (2004 May) Vol. 26, No. 4, pp. 295-318.

 Journal code: 7909595. ISSN: 0379-0355.
- CY Spain
- DT Bibliography
- LA English
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- EM 200501
- ED Entered STN: 21 Aug 2004

Last Updated on STN: 26 Jan 2005 Entered Medline: 25 Jan 2005

=> s L14 and cancer

536885 CANCER

L16 6 L14 AND CANCER

=> d L16 1-6 ti

L16 ANSWER 1 OF 6 MEDLINE on STN

TI Update on nonsteriodal anti-inflammatory drugs.

L16 ANSWER 2 OF 6 MEDLINE on STN

TI Gateways to clinical trials.

L16 ANSWER 3 OF 6 MEDLINE on STN

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L16 ANSWER 4 OF 6 MEDLINE on STN

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13951807 PY<2003

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L17 0 L16 AND PY<2003

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